

1 GTCTTCCACCATGCACTCGCTGGGCTTCTCTGTGGGTGTTCTCTGCTCGCCGCTG
 -----+-----+-----+-----+-----+-----+
 CAGGAAGGTGGTACGTAGCGACCCGGAAGAAGAGACACCGCACAAAGAGACGAGCGCGAC
 M H S L G F F S V A C S L L A A A - 60

61 CGCTGCTCCCGGGTCTCGGAGGCGCCGCGCGCGCGCTTCGAGTCCGGACTCG
 -----+-----+-----+-----+-----+-----+
 GCGAGAGGCGCCAGGAGCGCTCCGCGGGCGGGCGCGGCGGAAGCTCAGGCCTGAGC
 L L P G P R E A P A A A A A F E S G L D - 120

121 ACCTCTCGGACGCGGAGCCGACGCGGCGAGGCCACGGCTTATGCAAGCAAAGATCTGG
 -----+-----+-----+-----+-----+-----+
 TGGAGAGCCTGCGCCTCGGGCTGCGCCCGCTCCGGTGCCGAATACGTTCTTCTAGACC
 L S D A E P D A G E A T A Y A S K D L E - 180

181 AGGAGCAGTTACGGTCTGTGTCCAGTGTAGATGAACATGACTGTACTCTACCCAGAAT
 -----+-----+-----+-----+-----+-----+
 TCCTCGTCAATGCCAGACACAGGTCACATCTACTTGAGTACTGACATGAGATGGGTCTTA
 E Q L R S V S S V D E L M T V L Y P E Y - 240

241 ATTGGAATAATGTACAAGTGTACGCTAAGGAAGGAGGCTGGCAACATAACAGAGAACAGG
 -----+-----+-----+-----+-----+-----+
 TAACCTTTTACATGTTACAGTCGATTCCCTTCCCTCCGACCGTTGTATTGTCTCTGTCC
 W K M Y K C Q L R K G G W Q H N R E Q A - 300

CCAACCTCAACTCAAGGACAGAAGAGACTATAAAATTGCTGCAGCACATTATAATACAG

MATCH WITH FIG. 1B

FIG. 1A

MATCH WITH FIG. 1A

```

301  -----+-----+-----+-----+-----+-----+-----+-----+-----+-----+
      GGTGGAGTTGAGTTCCTCTCTCTGATATTTTAAACGACGTCGTTGTAATATATGTC
          N L N S R T E E T I K F A A A H Y N T E -
360

361  -----+-----+-----+-----+-----+-----+-----+-----+-----+-----+
      AGATCTTGAAAGTATGATAATGAGTGGAGAAAGACTCAATGCATGCCACGGGAGGTGT
          I L K S I D N E W R K T Q C M P R E V C -
420

421  -----+-----+-----+-----+-----+-----+-----+-----+-----+-----+
      GTATAGATGTGGGAAGGAGTTTGGAGTCGCGACAAACACCTTCTTTAAACCTCCATGTG
          C A T A T C A C A C C C T T C C T C A A A C C T C A G C G T G T T G T G G A A G A A A T T T G G A G G T A C A C
          I D V G G K E F G V A T N T F F K P P C V -
480

481  -----+-----+-----+-----+-----+-----+-----+-----+-----+-----+
      TGTCCGCTACAGATGTGGGGGTTGCTGCAATAGTGAGGGGTCAGTGCAATGAACACCA
          A C A G G C A G A T G T C T A C A C C C C A A C G A C G T T A T C A C T C C C G A C G T C A C G T A C T T G T G G T
          S V Y R C G G C C N S E G L Q C M N T S -
540

541  -----+-----+-----+-----+-----+-----+-----+-----+-----+-----+
      GCACGAGTACCTCAGCAAGACGTTATTGAAATTACAGTGCCCTCTCTCTCAAGGCCCA
          C G T G C T C G A T G G A G T C G T T C T G C A A T A A A C T T T A A T G T C A C G G A G A G A G T T C C G G G T
          T S Y L S K T L F E I T V P L S Q G P K -
600

601  -----+-----+-----+-----+-----+-----+-----+-----+-----+-----+
      AACCAGTAAACAATCAGTTTGGCCAATCACACTTCCTGCCGATGCATGTCTAAACTGGATG
          T T G G T C A T T G T T A G T C A A A C G G T T A G T G T G A A G G A C G G T A C G T A C A G A T T T G A C C T A C
          P V T I S F A N H T S C R C M S K L D V -
660

```

MATCH WITH FIG. 1C

FIG. 1B

MATCH WITH FIG. 1B

TTTACAGACAAGTTCATTCCATTATTAGACGTTCCCTGCCAGCACTACCACAGTGC
661-----+-----+-----+-----+-----+-----+-----+ 720

AAATGCTCTGTTCAAGTAAGGTAATAATCTGCAAGGACGGTCGTTGTGATGGTGTCAACAG
Y R Q V H S I I R R S L P A T L P Q C Q

AGGCAGCGAACAGACCCTGCCCCACCAATTACATGTGGAATAATCACATCTGCAGATGCC
721 -----+-----+-----+-----+-----+-----+-----+ 780

TCCGTCGCTTGTTCTGGACGGGGTGGTTAATGTACACCTTATTAGTGTAGACGCTCTACGG
A A N K T C P T N Y M W N N H I C R C L

781 TGGCTCAGGAAGATTTTATGTTTCCCGGATGCTGGAGATGACTCAACAGATGGATTCC 840

ACCGAGTCCTCTAAATAACAAAGGAGCCTACGACCTCTACTGAGTTGTCTACCTAAGG
A Q E D F M F S S D A G D D S T D G F H

841 ATGACATCTGTGGACCAACAAGGAGCTGGATGAAGAGACCCTGTCACTGTGTCTGCAGAG
-----+-----+-----+-----+-----+-----+-----+ 900

TACTGTAGACACCTGGTTTGTTCTCGACCTACTTCTCTGGACAGTCACACAGAGCTCTC
D I C G P N K E L D E E T C O C V C R A -

CGGGGCTTCGGCCCTGCCAGCTGTGGACCCCAAGAACTAGACAGAACTCATGCCAGT
 901 -----+-----+-----+-----+-----+-----+ 960

G C C C C G A A G C C G G A C G G T C G A C A C C T G G G G T G T T C T T G A T C T G T C T T T G A G T A C G G T C A
 G L R P A S C G P H K E L D R N S C O C -

GTGCTGTAAAAACAACACTCTTCCCCAGCCCAATGTGGGCCAACCGAGAATTGATGAAA
-----+-----+-----+-----+-----+-----+-----+-----+-----+-----+
961

CACAGACATTTTGTGAGAGGGTCCGTTACACCCGGTTGGCTCTTAAACTACTTT

MATCH WITH FIG. 1D

FIG. 1C

MATCH WITH FIG. 1C

V	C	K	N	K	L	F	P	S	Q	C	G	A	N	R	E	F	D	E	N	-
AC	AT	GC	CC	AG	TG	TAT	G	TAA	AG	AA	CC	TG	CC	CC	AG	AA	TC	AA	TC	CT
1021	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---
	TG	TG	TA	CG	GT	C	A	C	A	T	AT	TT	CT	TG	GC	GG	GT	CT	TT	AG
	T	C	Q	C	V	C	K	R	T	C	P	R	N	Q	P	L	N	P	G	K
	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---
	AA	TG	TC	CT	GT	GA	AT	GT	AC	AG	AA	GT	CC	AC	AG	AA	AT	GT	TC	CT
1081	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---
	TT	AC	CG	GA	CA	CT	T	AC	AT	GT	CT	TT	AC	GA	CA	AA	TT	TC	CT	TT
	C	A	C	E	C	T	E	S	P	Q	K	C	L	L	K	G	K	K	F	H
	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---
	AC	CA	AA	CA	TC	GC	AG	CT	GT	T	AC	AG	AC	GG	CC	AT	GT	GC	AG	AG
1141	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---
	TG	GT	GG	TT	TG	TA	CG	TC	GA	CA	AT	GT	CT	GC	CG	GT	AC	AT	GT	CT
	H	Q	T	C	S	C	Y	R	R	P	C	T	N	R	Q	K	A	C	E	P
	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---
	CA	GG	AT	TT	TC	AT	AT	AG	AG	AG	TC	GT	GT	GT	CC	CT	TC	AT	AT	TC
1201	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---
	GT	CT	AA	AG	T	AT	CA	CT	TC	TC	AC	AG	CA	CA	AG	GA	AG	T	AT	AA
	G	F	S	Y	S	E	E	V	C	R	C	V	P	S	Y	W	Q	R	P	Q
	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---
	AA	AT	GA	GC	TA	AG	AT	GT	FA	CT	GT	TT	TC	CA	GT	TC	GA	TT	TC	TA

MATCH WITH FIG. 1E

FIG. 1D

[illegible]

TTTACTCGATTCTAACATGACAAAGGTCAAGTAGCTAAAGATAATACCTTTTGACACA

*
S
M

1321 TGCCACAGTAGAACTGTCTGTGAACAGAGAGACCCCTTGTGGGTCCATGCTAACAAAGACA
-----+-----+-----+-----+-----+-----+ 1380

ACGGTGTCACTTGACAGACACTTGTCTCTCTGGGAACCCAGGTACGATGTTTCTGT

1381 AAAGTCTGCTTTCCCTGAACCATGTGGATAACTTTACAGAAATGGACTGGAGCTCATCTG 1440

TTTCAGACAGAAAGGACTTGGTACACCTATTGAAATGTCTTTACCTGACCTCGAGTAGAC

1441 CAAAGGCTCTTGTAAGACTGGTTTCTGCCAATGACCAACAGATTTCCTC
-----+-----+-----+-----+-----+-----+ 1500
GTTTCCGGAGAACATTCTTGACCAAAAGACGGTACTGGTTTGTCTGTTCTAAAGGAG

TTGTGATTCTTTAAAGAATGACTATATAATTTATTTCCTAAATAATGTTCTGC
- - - - - + - - - - - + - - - - - + - - - - - + - - - - - + - - - - - + 1501
AACACTAAAGAAATTTCTTACTGATATATTAATAAAGGTGATTTTATATAACAAAGACG

561
ATTCATTTTATAGCAACAACATTGGTAAACTCACTGTGATCAATATTTTATATCAT
-----+-----+-----+-----+-----+-----+ 1620
TAAGTAAAAAATATCGTTGTTGTTAACCATTTTGAGTGACACTAGTTATAAAAAATATAGTA

GCAAAATATGTTAAATAAATGAAAATTGTATTTATATAAAAAA
621 -----+-----+-----+-----+-----
CGTTTTATACAAATTTTATTTACTTTTAACATAAATTTTTTTTTT

FIG. 1E

```

1  CGAGGCCACGGCTTATGCAAGCAAGATCTGGAGGAGCAGTTACGGTCTGTGTCCAGTGT
-----+-----+-----+-----+-----+-----+-----+-----+
71 AGATGAACATGACTGTACTCTACCCAGAATAATTGGAAAATGTACAAGTGTACGCTAAG
-----+-----+-----+-----+-----+-----+-----+-----+
      M T V L Y P E Y W K M Y K C Q L R
121 GAAAGGAGGCTGGCAACATAACAGAGAACAGGCCAACCTCAACTCAAGGACAGAAGAGAC
-----+-----+-----+-----+-----+-----+-----+-----+
      K G G W Q H N R E Q A N L N S R T E E T
181 TATAAAATTGCTGCACACATTAATAACAGAGATCTTGAAAGTATTGATAATGAGTG
-----+-----+-----+-----+-----+-----+-----+-----+
      I K F A A A H Y N T E I L K S I D N E W
241 GAGAAAGACTCAATGCATGCCACGGGAGGTGTATAGATGTGGGAAGGAGTTTGGAGT
-----+-----+-----+-----+-----+-----+-----+-----+
      R K T Q C M P R E V C I D V G K E F G V
301 CGCGACAAACACCTTCTTTAAACCTCCATGTGTGTCCTACAGATGTGGGGTTCCTG
-----+-----+-----+-----+-----+-----+-----+-----+
      A T N T F F K P P C V S V Y R C G G C C

```

FIG. 2A

361 CAATAGTGGGCTGCAGTGCATGAACACGACGAGCTACCTCAGCAAGACGTTATT
 N S E G L Q C M N T S T S Y L S K T L F
 421 TGAAATTACAGTGCCTCTCTCTCAAGGCCCAACGAGTAACAATCAGTTTGTGCCAATCA
 E I T V P L S Q G P K P V T I S F A N H
 481 CACTTCCTGCCGATGCATGTCTAACTGGATGTTACAGACAAGTTCATTCCATTATTAG
 T S C R C M S K L D V Y R Q V H S I I R
 541 ACGTTCCCTGCCAGCAACACTACCACAGTGTGAGGAGCGAACAAGACCTGCCCCACCAA
 R S L P A T L P Q C Q A A N K T C P T N
 601 TTACATGTGGAATAATCACATCTGCAGATGCCCTGGCTCAGGAAGATTTATGTTTCCTC
 Y M W N N H I C R C L A Q E D F M F S S
 661 GGATGCTGGAGATGACTCAACAGATGGATTCCATGACATCTGTGGACCAACAAGGAGCT
 D A G D D S T D G F H D I C G P N K E L

FIG. 2B

```

721  GGATGAAGAGACCTGTCAGTGTGTCTGCAGAGCGGGCTTCGGCCCTGCCAGCTGTGGACC
      -----+-----+-----+-----+-----+-----+-----+
      D E E T C Q C V C R A G L R P A S C G P

781  CCACAAAGAACTAGACAGAACTCATGCCAGTGTGTCTGTAAACAACTCTTCCCCAG
      -----+-----+-----+-----+-----+-----+-----+
      H K E L D R N S C Q C V C K N K L F P S

841  CCAATGTGGGCCCAACCGAGAAATTGATGAAACACATGCCAGTGTGTATGTAAAGAAC
      -----+-----+-----+-----+-----+-----+-----+
      Q C G A N R E F D E N T C Q C V C C K R T

901  CTGCCCCAGAAATCAACCCCTAAATCCTGGAAATGTGCCCTGTGAATGTACAGAAAGTCC
      -----+-----+-----+-----+-----+-----+-----+
      C P R N Q P L N P G K C A C E C T E S P

961  ACAGAAATGCTTGTTAAAGGAAGAAGTTCACCACCAACATGCAGCTGTACAGACG
      -----+-----+-----+-----+-----+-----+-----+
      Q K C L L K G K K F H H Q T C S C Y R R

1021 GCCATGTACGAACCGCAGAGGCTTGTGAGCCAGGATTTTCATATAGTGAAGAAGTGTG
      -----+-----+-----+-----+-----+-----+-----+
      P C T N R Q K A C E P G F S Y S E E V C

```

FIG. 2C

FIG. 2D

FIG. 2D

FIG. 3A

FIG. 3A

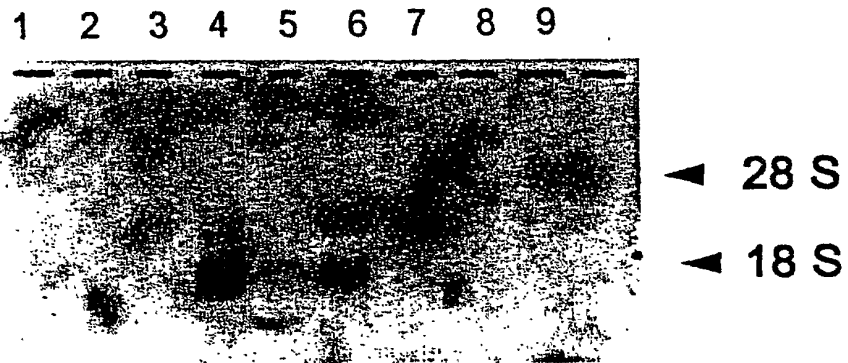
201 250
Pdgfa TSLNPD YREEDTDVR. GKHKFKH TH DKTALKETLG
Pdgfb RSPGGSQEQR AKTPQTRVTI RTVRVRRPPK KSRYSWSVY VGARCCCLMPW SLPGPHP ...
Vegf RCK..... GKQKRRK KSRYSWSVY VGARCCCLMPW SLPGPHP ...
Vegf2 RRSLPATLPQ COAANKTCPT NYMNNHICR CLAQEDFMFS SDAGDDSTDG
251 300
Pdgfa
Pdgfb A..... CSE RRKHLFVQDP QTCKCCKNT
Vegf CGP..... RAGLRPASC G PHKEL... DR NSCQCVCCKNK
Vegf2 FHDICGPNKE LDEETCQCVC VCKRTCPRNQ PLNPGKCACE CTESPKKCLL
301 350
Pdgfa
Pdgfb LELNERTCRC DKPRR.....
Vegf ..DSRCKARQ LELNERTCRC DKPRR.....
Vegf2 LFPSQCGANR .EFDENTCQC VCKRTCPRNQ PLNPGKCACE CTESPKKCLL
351 398
Pdgfa
Pdgfb
Vegf SCYRRPCTNR QKACEPGFSY SEEVCRCPVS YWQRPQMS
Vegf2 KGKKFHHQTC SCYRRPCTNR QKACEPGFSY SEEVCRCPVS YWQRPQMS

FIG. 3B

| PERCENTAGE (%) OF AMINO ACID IDENTITIES BETWEEN
EACH PAIR OF GENES IS SHOWN IN THE
FOLLOWING TABLE | | | | |
|--|---------------|--------------|------|-------|
| | PDGF α | PDGF β | VEGF | VEGF2 |
| PDGF α | | | | |
| PDGF β | 48.0 | | | |
| VEGF | 20.7 | 22.7 | | |
| VEGF2 | 23.5 | 22.4 | 30.0 | |

FIG. 4

**Expression of VEGF2 mRNA in
Human Breast Tumor Cells**

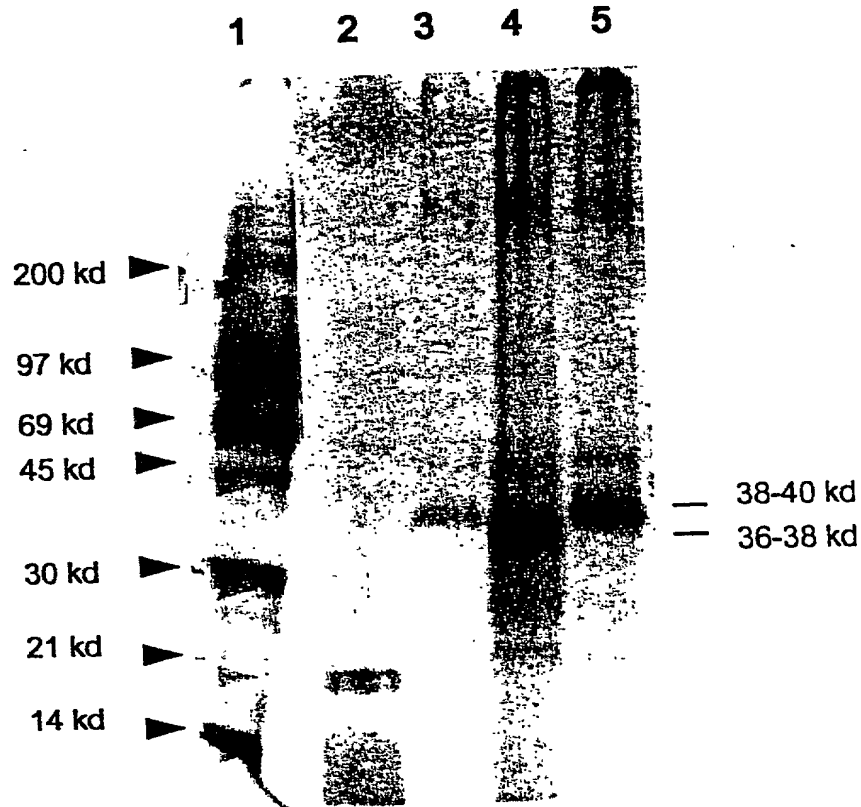


Lane 1. normal breast tissue
Lane 2. breast tumor tissue
Lane 3-9. breast tumor cell lines.

FIG.5



FIG. 6



Lane 1: 14-C and rainbow M.W. marker
 Lane 2: FGF control
 Lane 3: VEGF2 (M13-reverse & forward primers)
 Lane 4: VEGF2 (M13-reverse & VEGF-F4 primers)
 Lane 5: VEGF2 (M13-reverse & VEGF-F5 primers)

FIG.7

non-reducing gel

M 1 M 2 M

98 —
67 —
45 —
30 —
21 —
14 —

Lane M
Lane 1
Lane 2

Marker
vector medium
VEGF2 medium

FIG.8A

reducing gel

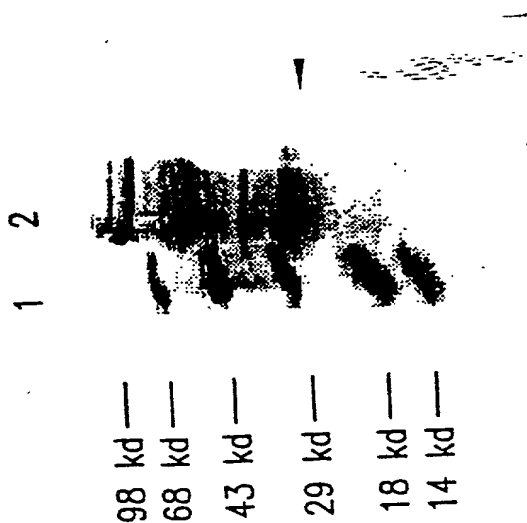
M 1 2 M 3 4 M

210 —
98 —
67 —
45 —
30 —
21 —
14 —

Lane M:
Lane 1:
Lane 2:
Lane 3:
Lane 4:

Marker
vector Cytoplasm
vector medium
VEGF2 Cytoplasm
VEGF2 medium

FIG.8B



Lane 1: Molecular weight marker
Lane 2: Precipitates containing VEGF2.

FIG.9

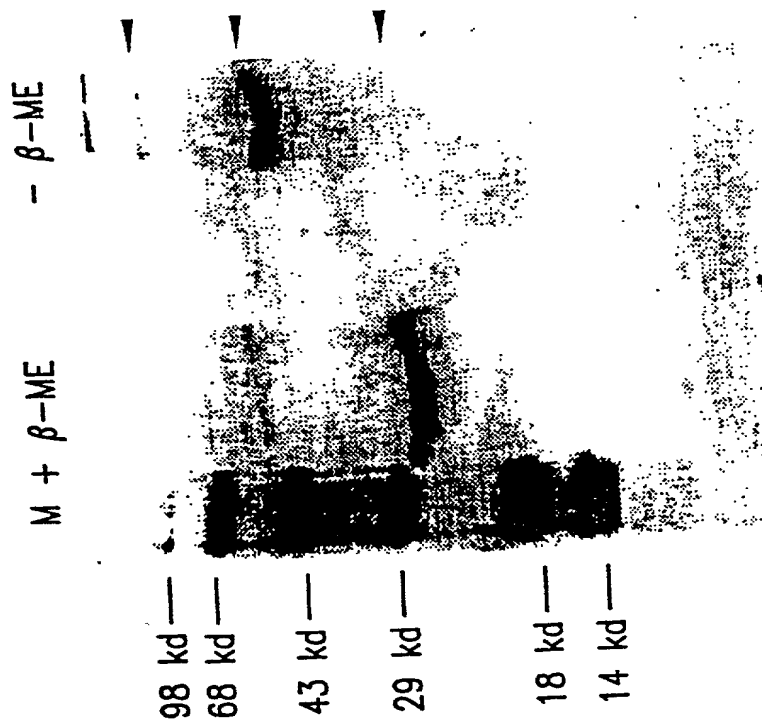


FIG.10

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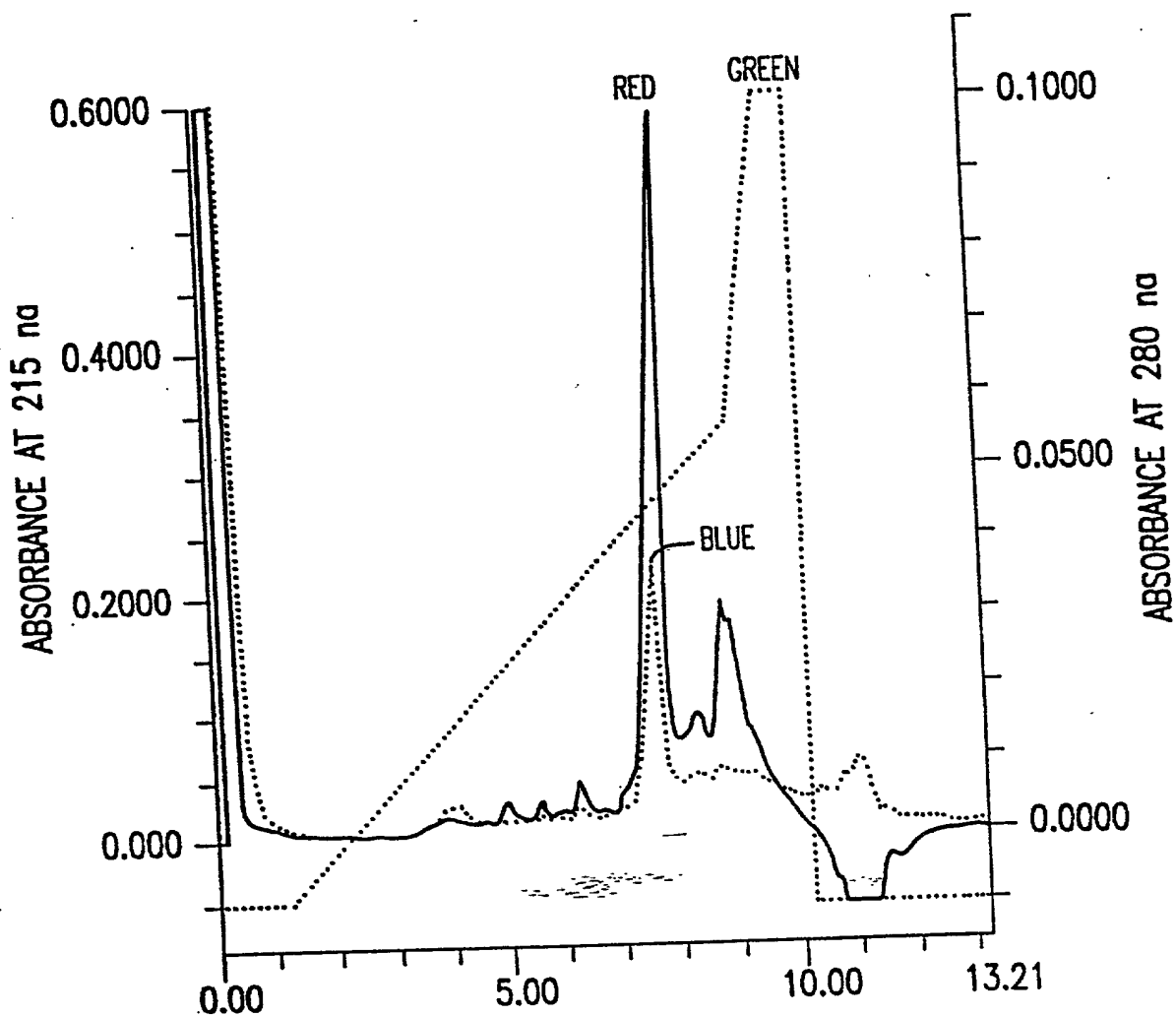


FIG. 11

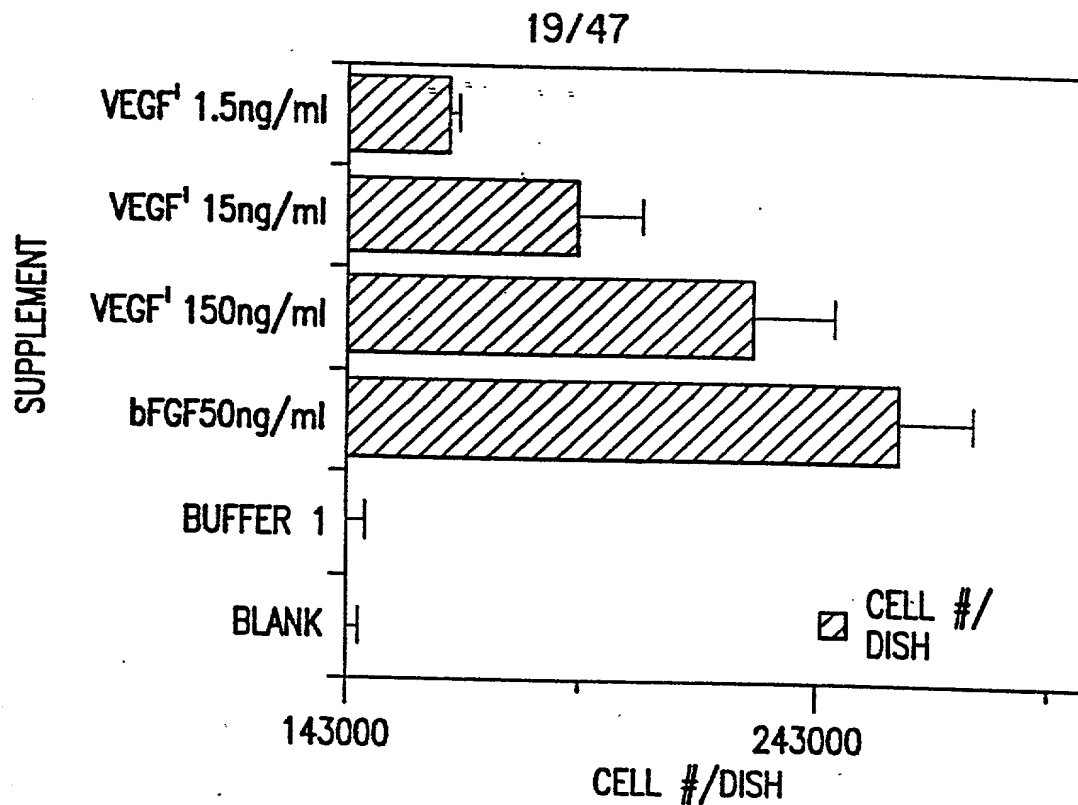


FIG.12

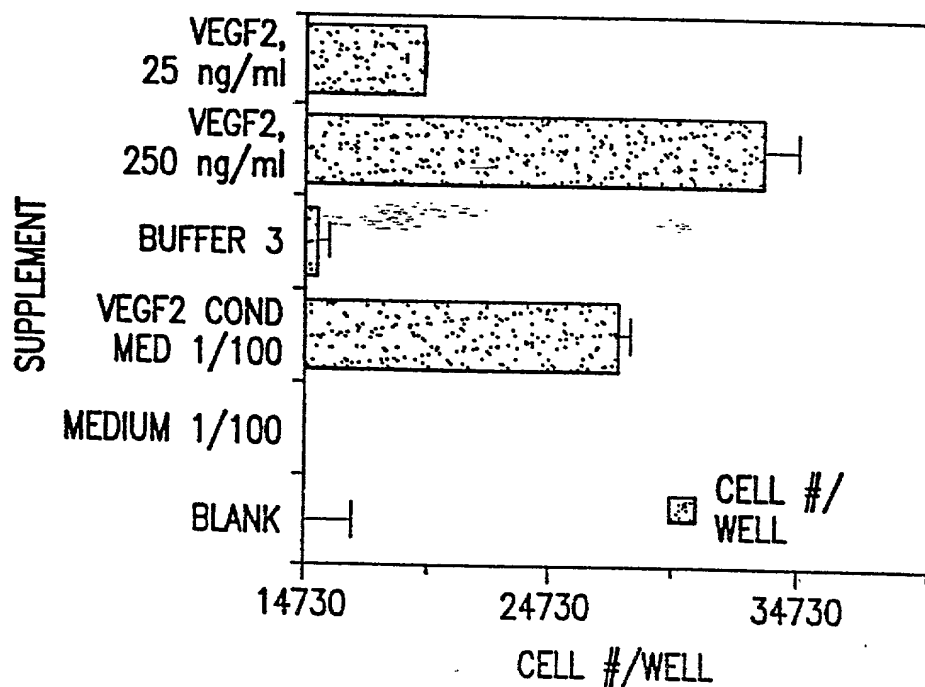


FIG.13

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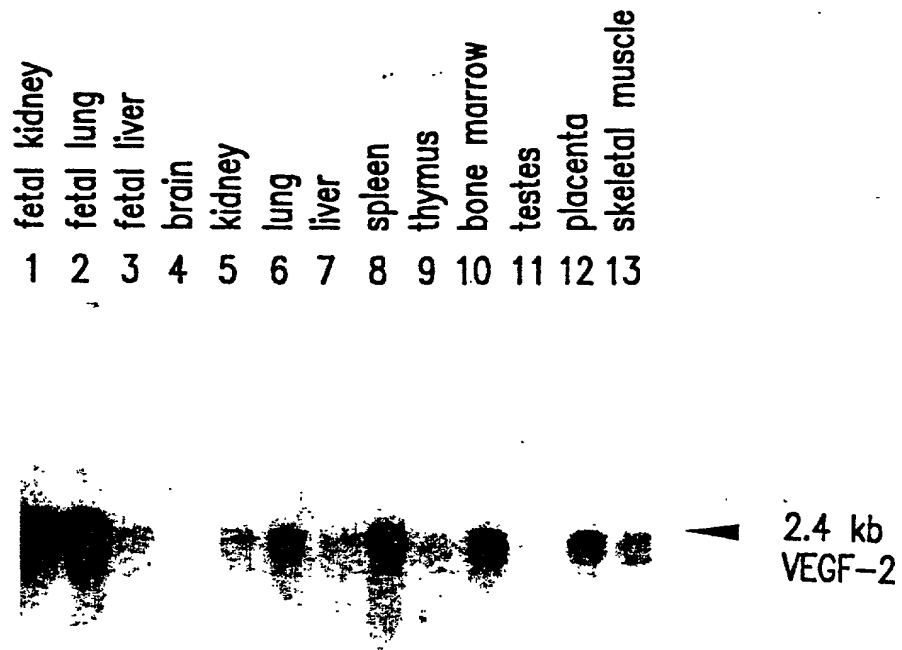


FIG.14A

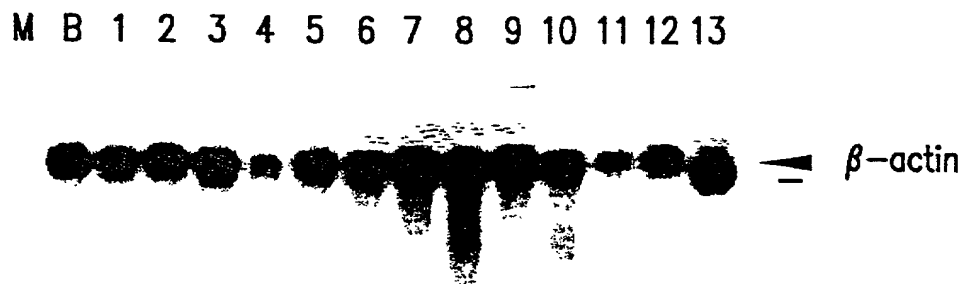
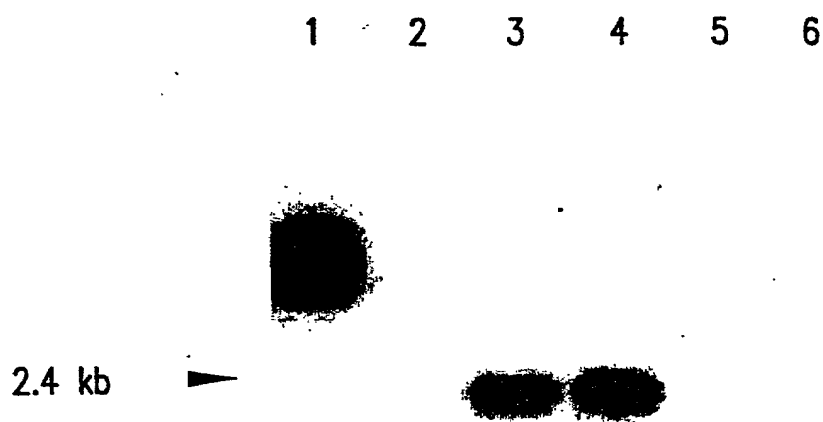


FIG.14B

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1. Molecular Weight Marker
2. umbelical vein endothelial cells
3. aortic smooth muscle cells
4. Dermal fibroblast

FIG.15

1. m.w. marker
2. blank
3. control protein-HA
4. vector control
5. VEGF2-HA

96 kd —
68 kd —
45 kd —
30 kd —
21 kd —
14.3 kd —

conditioned medium

FIG.16A

1. m.w. marker
2. blank
3. control protein-HA
4. VEGF2-HA
5. vector control

53 kd —
68 kd —
96 kd —
45 kd —
30 kd —
21 kd —
14.3 kd —
54 kd —

cell lysate
FIG.16B

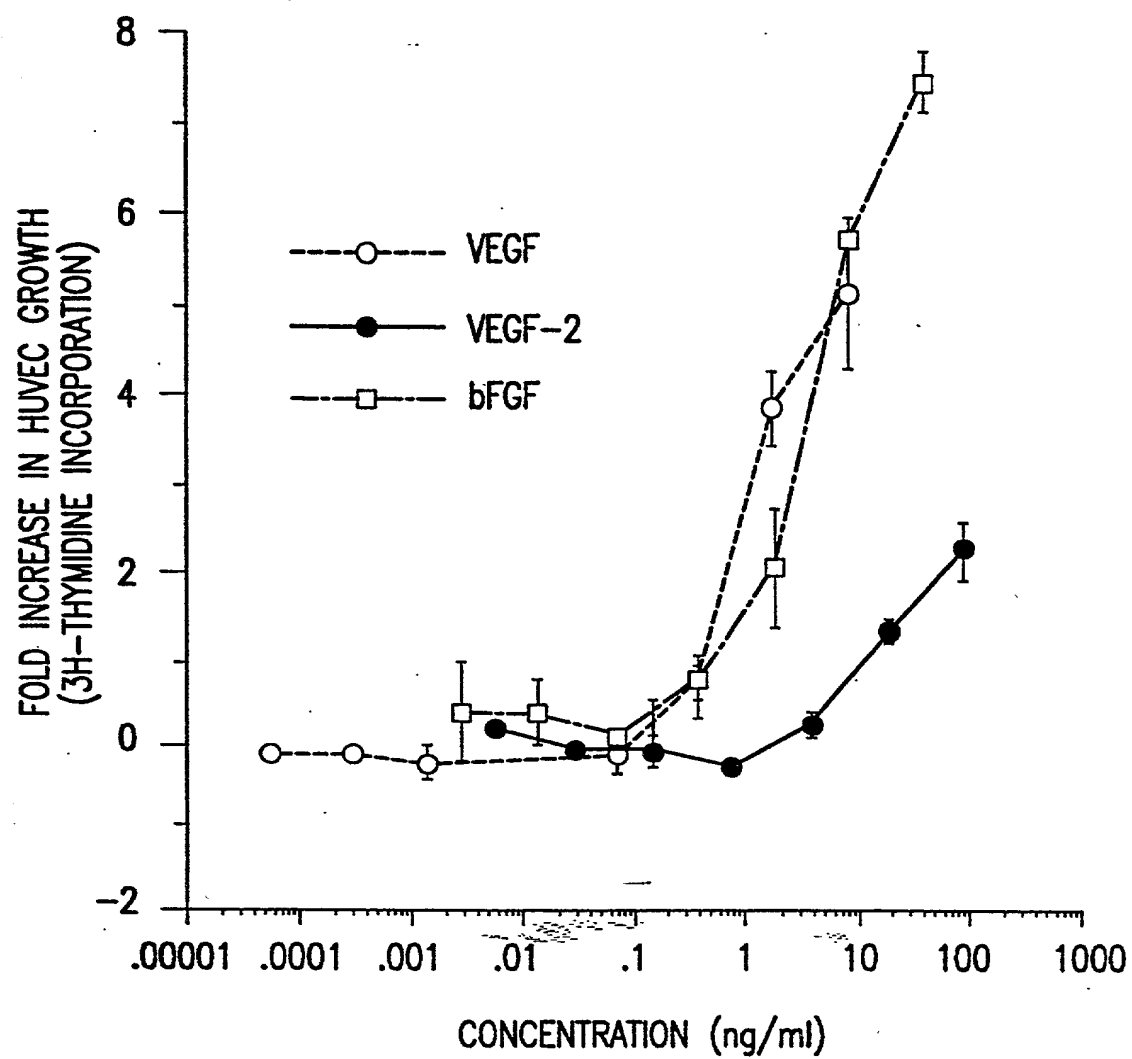


FIG.17

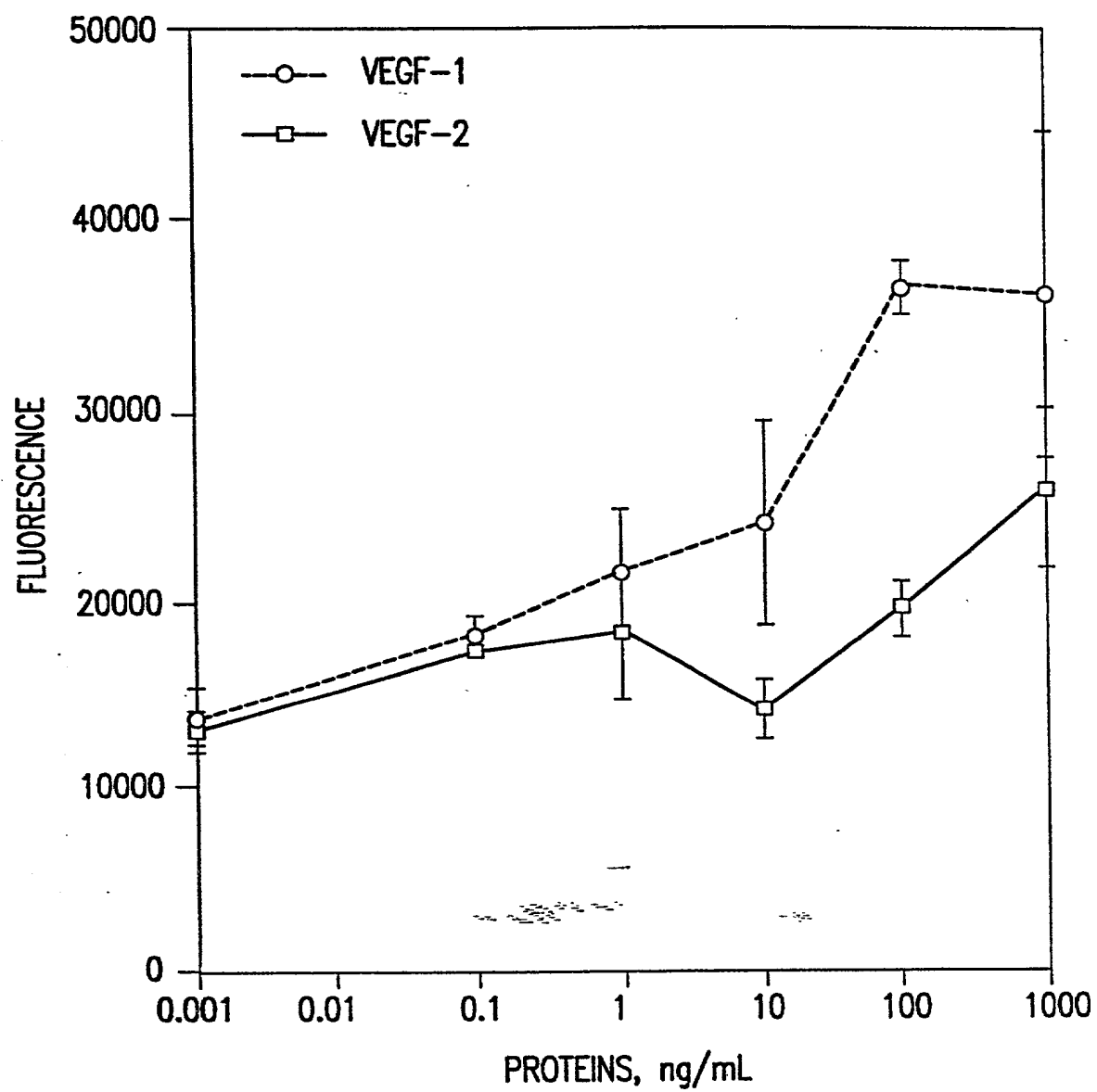


FIG.18

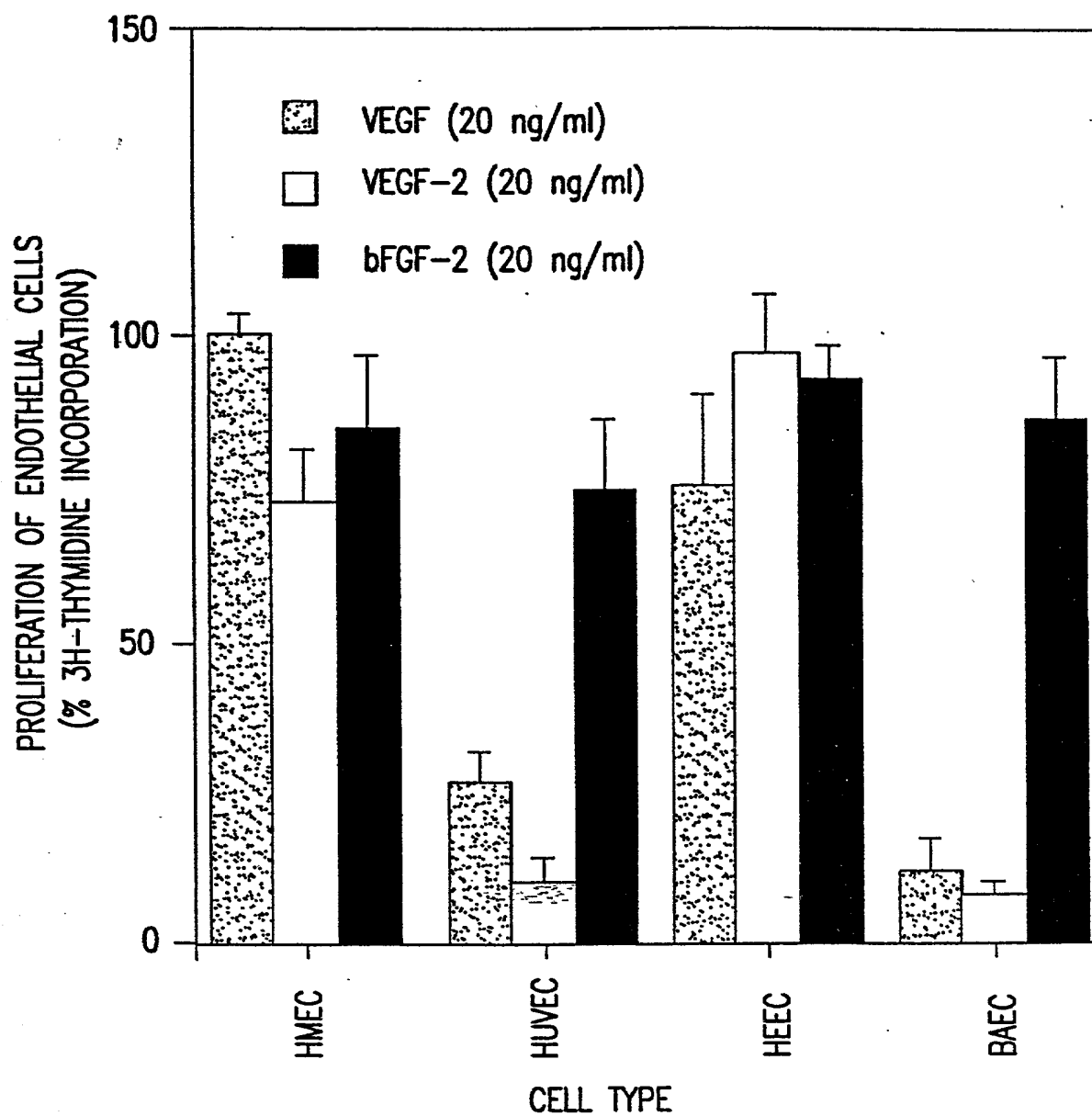


FIG.19

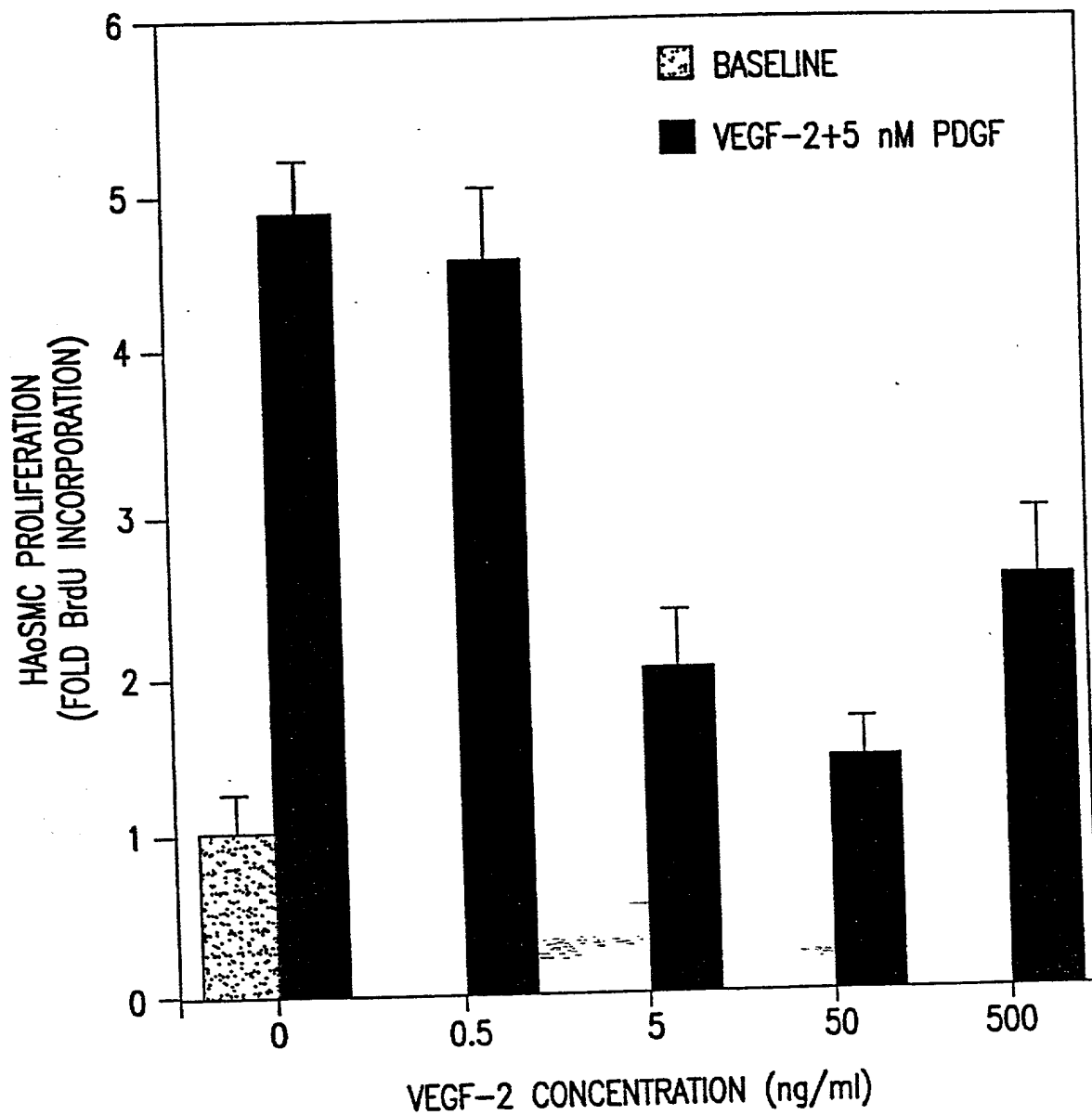


FIG.20A

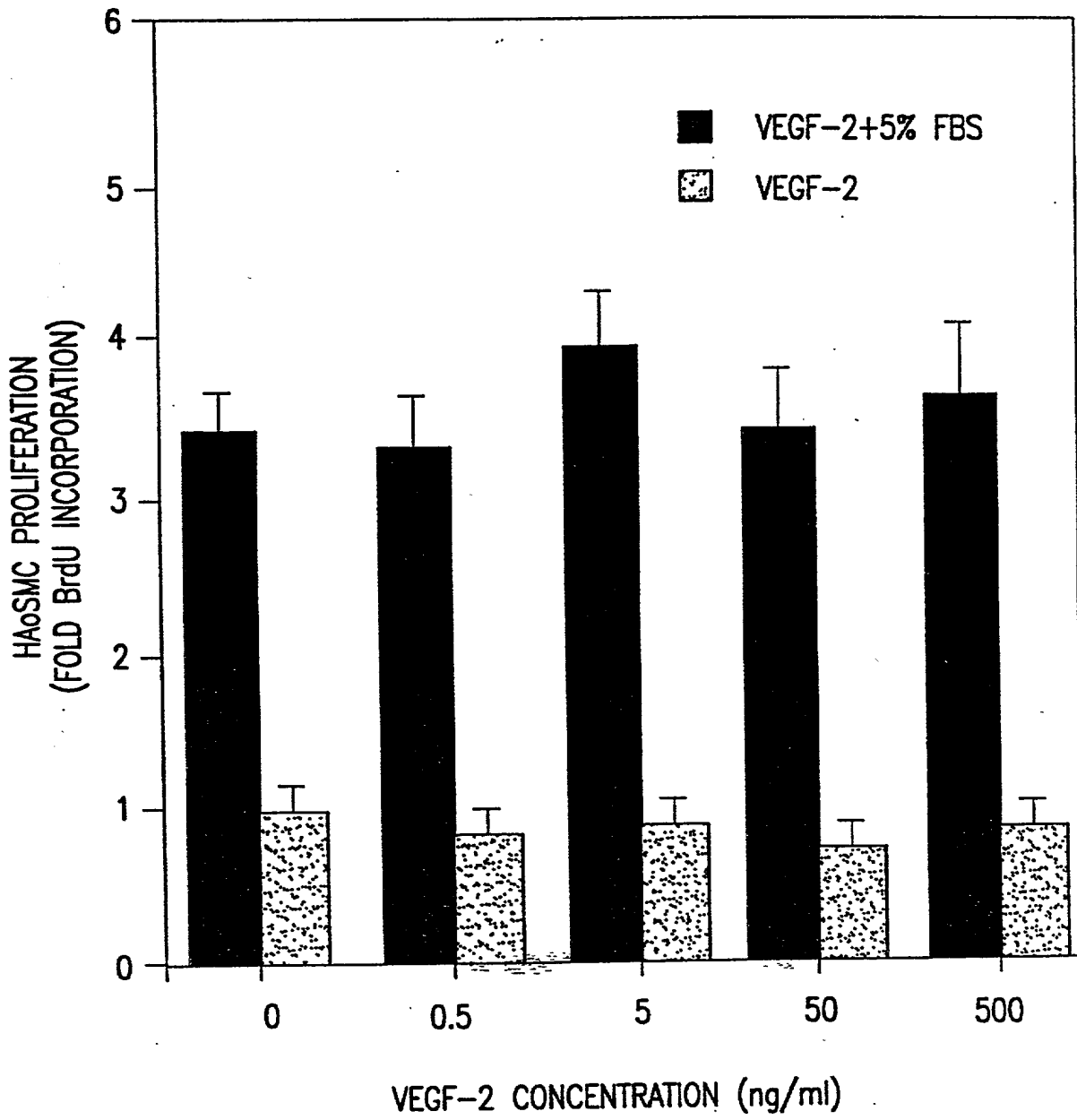


FIG.20B

HUVEC MIGRATION

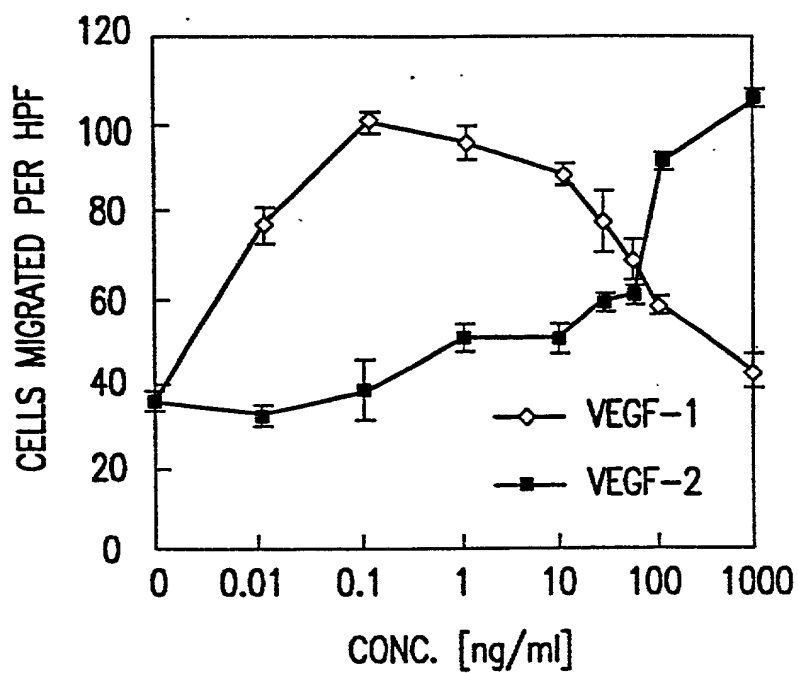


FIG.21A

BMEC MIGRATION

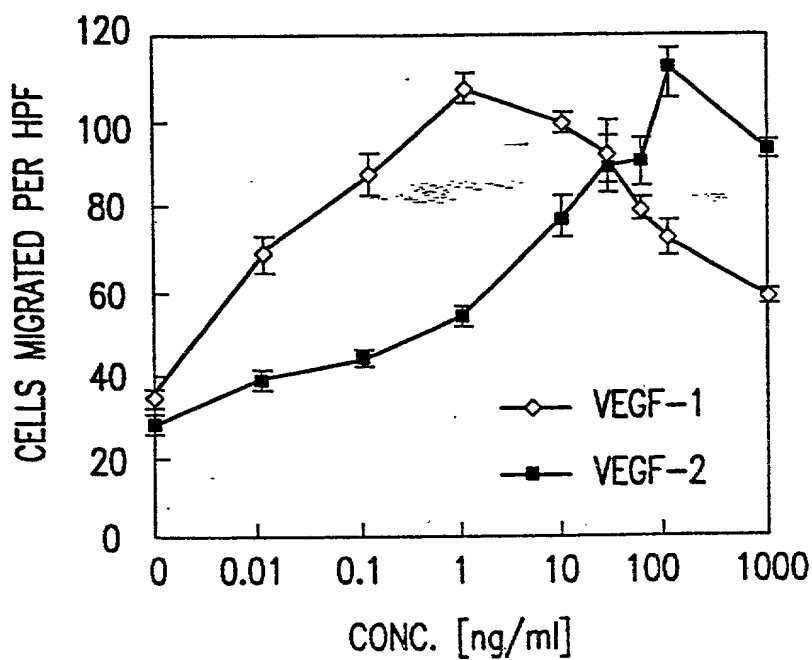


FIG.21B

HUVEC-NO-RELEASE

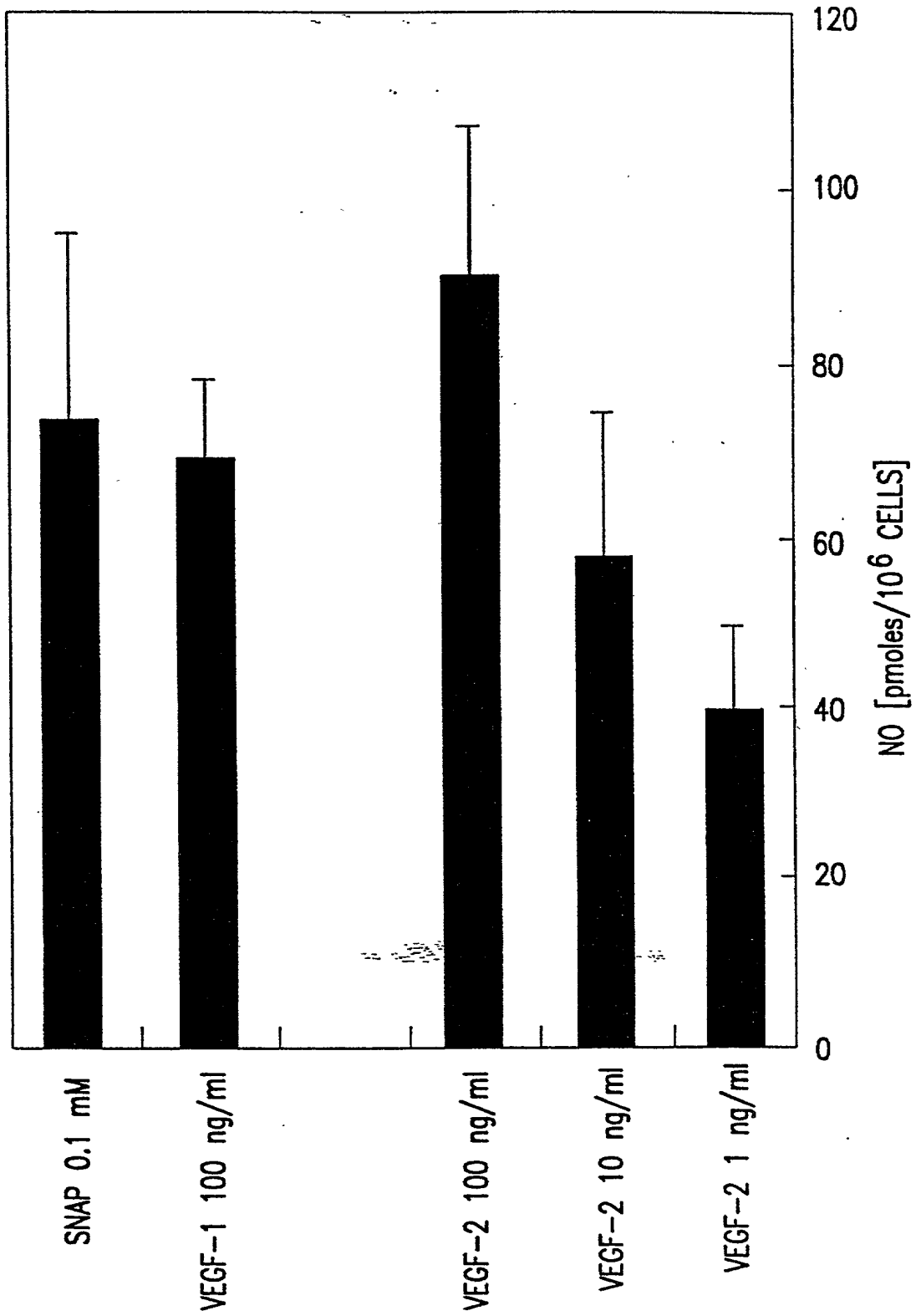


FIG.22

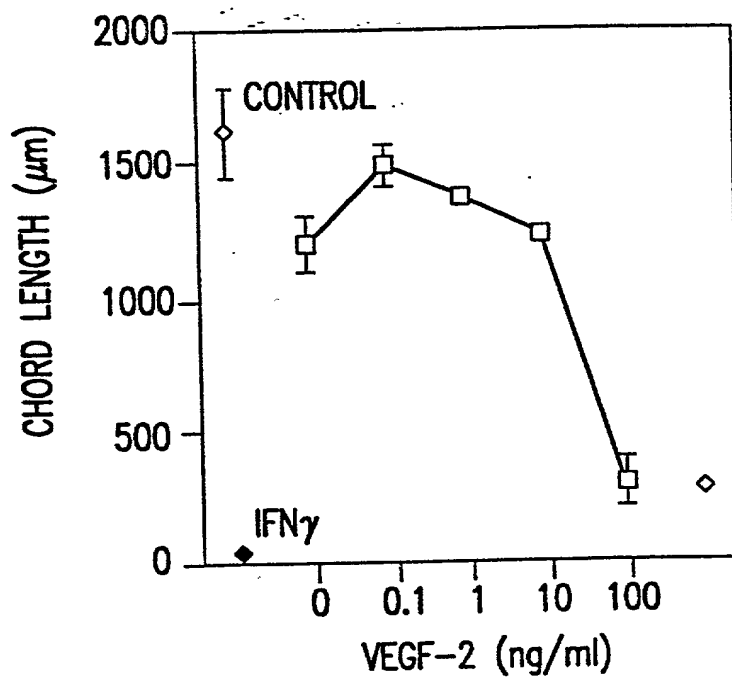


FIG.23

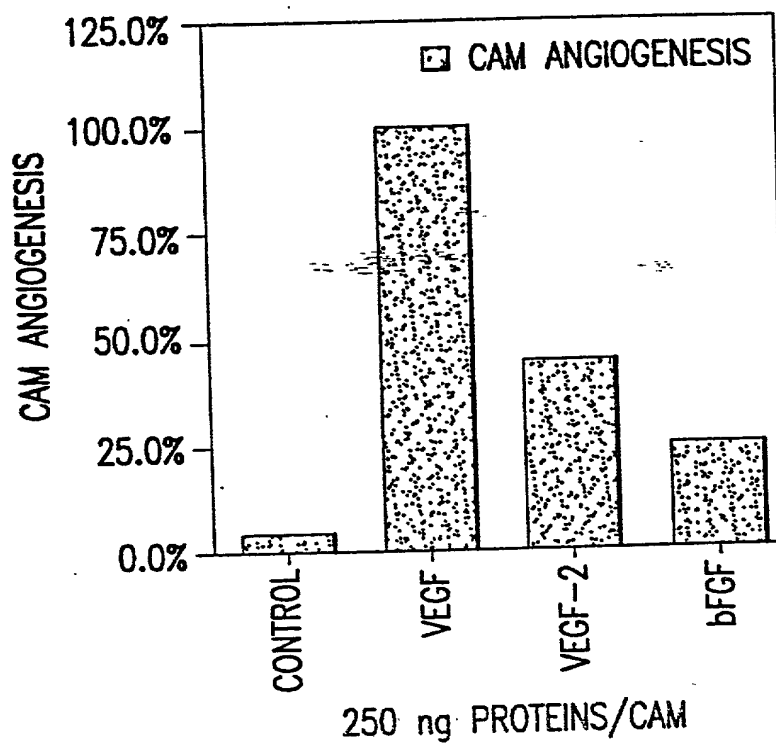


FIG.24

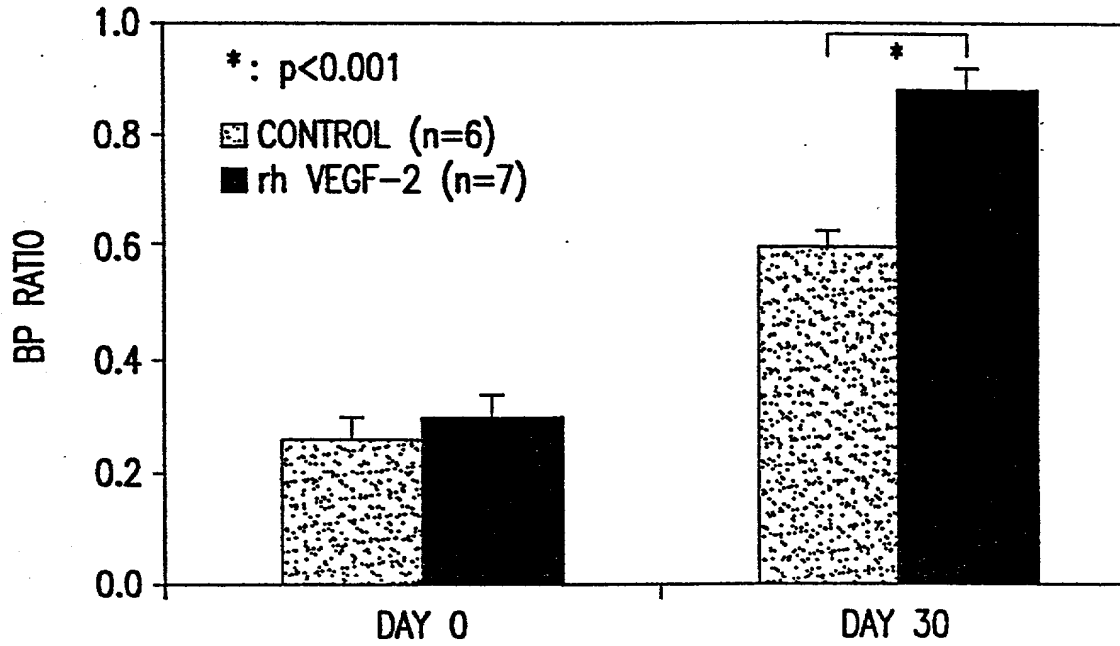
CALF BLOOD PRESSURE RATIO
-PROTEIN I.A.-

FIG.25A

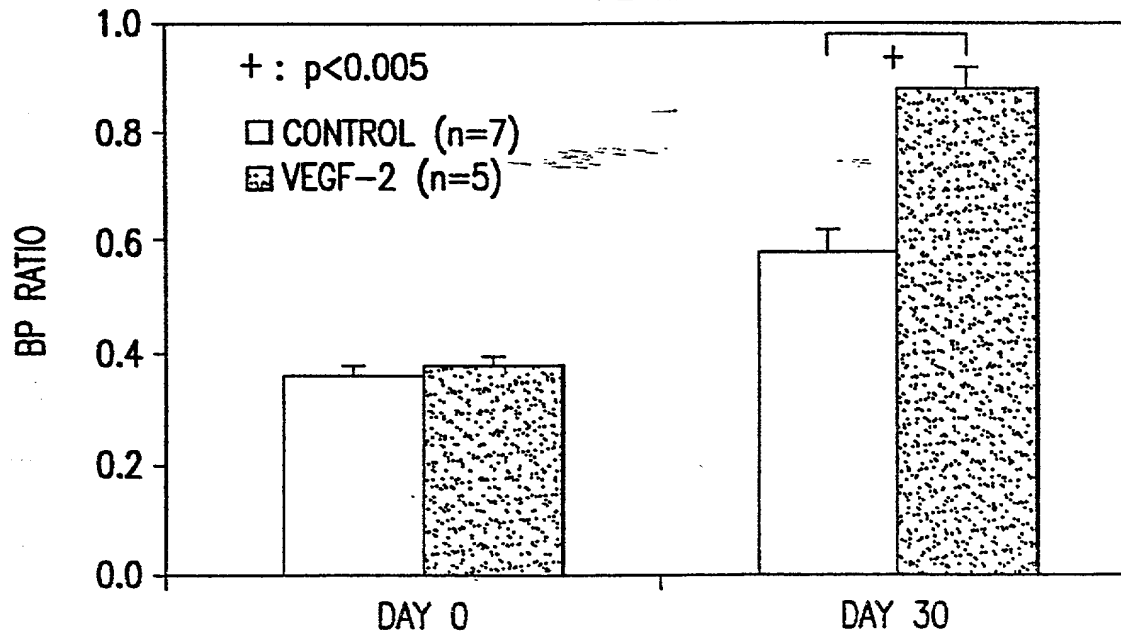
CALF BLOOD PRESSURE RATIO
-PLASMID-

FIG.25B

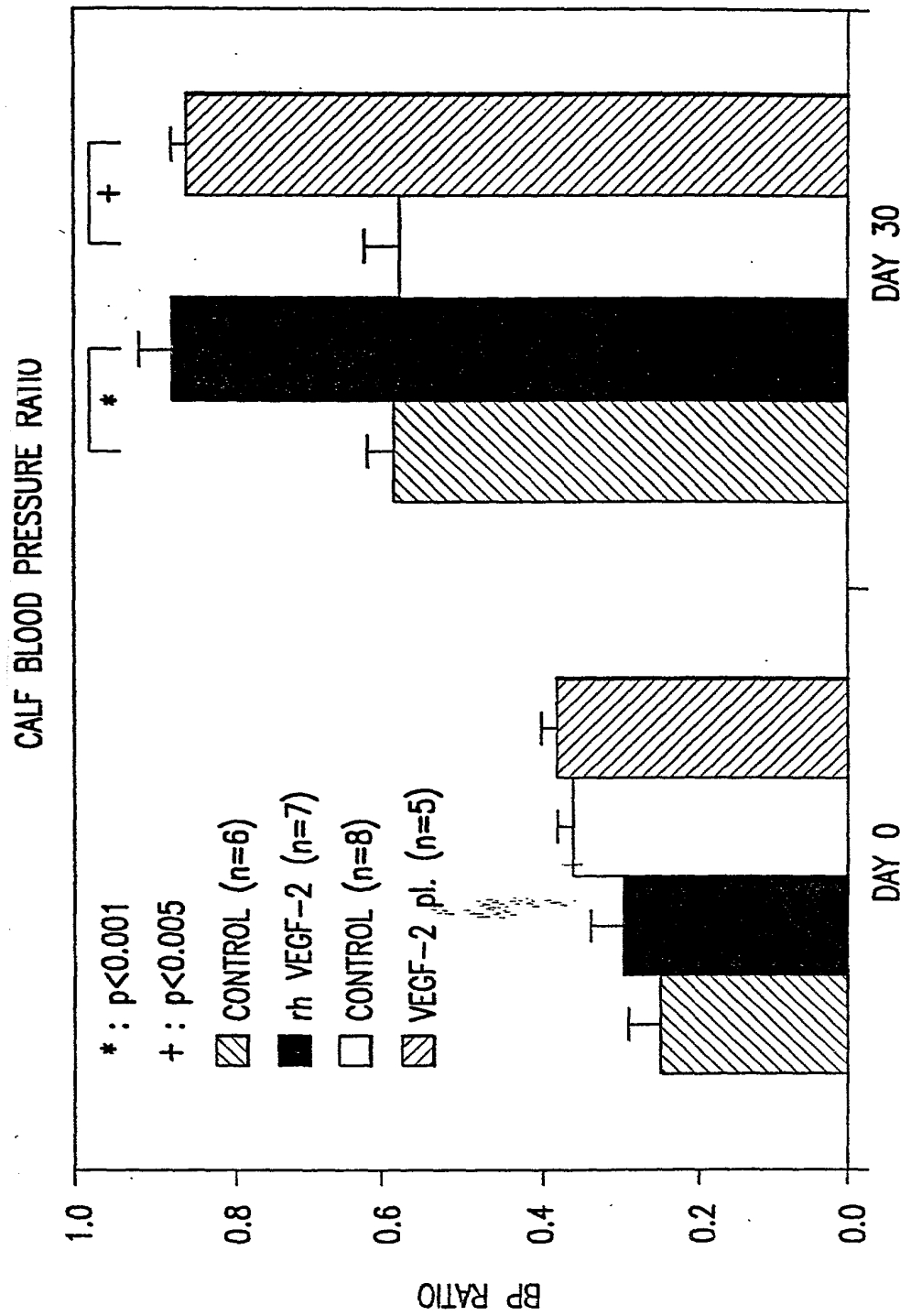


FIG.25C

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LLIAC BLOOD FLOW
-PROTEIN I.A.-

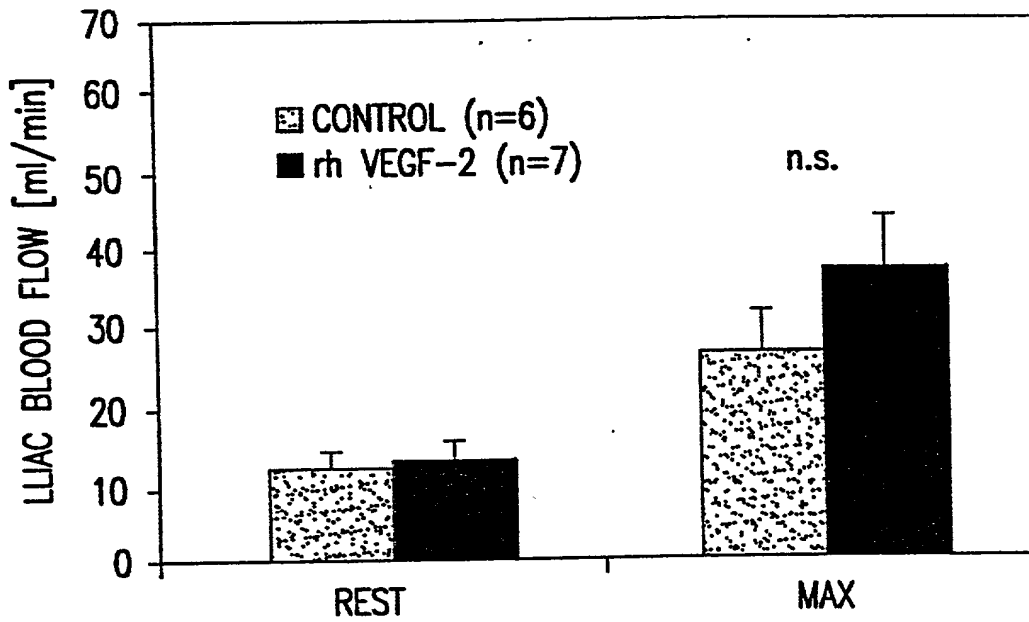


FIG.25D

LLIAC FLOW RESERVE
-PROTEIN I.A.-

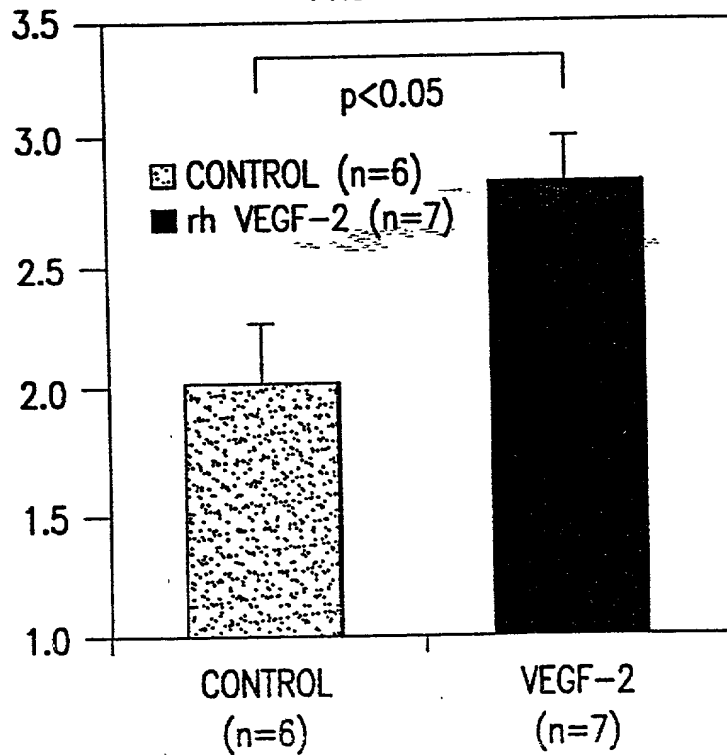


FIG.25E

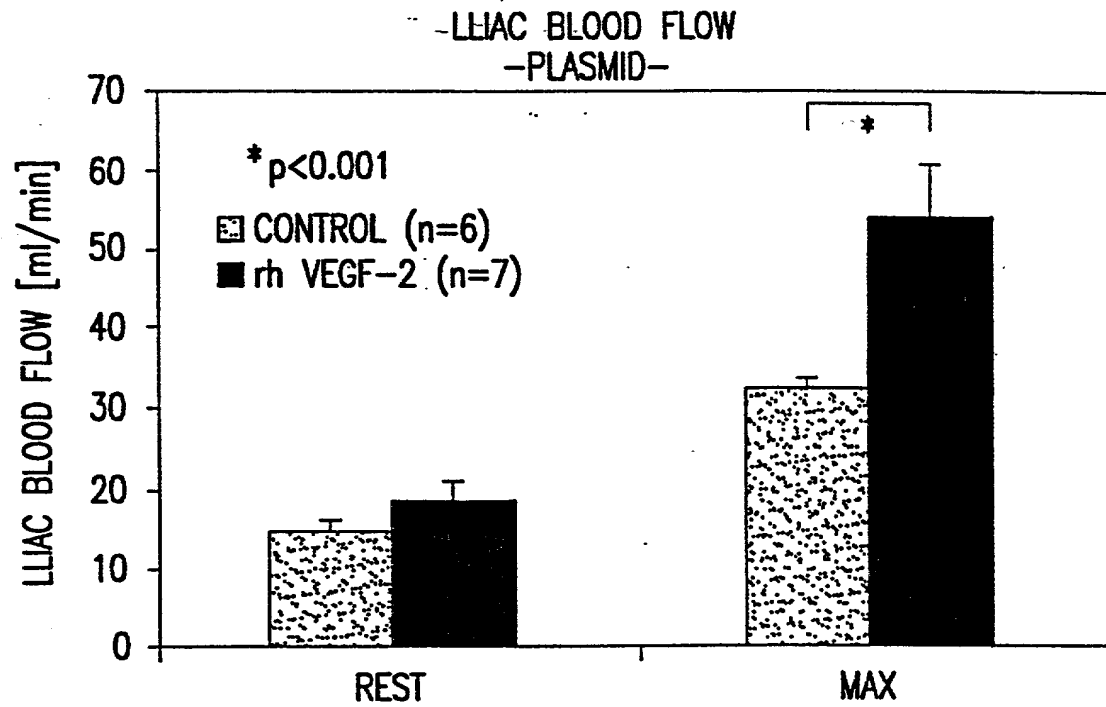


FIG.25F

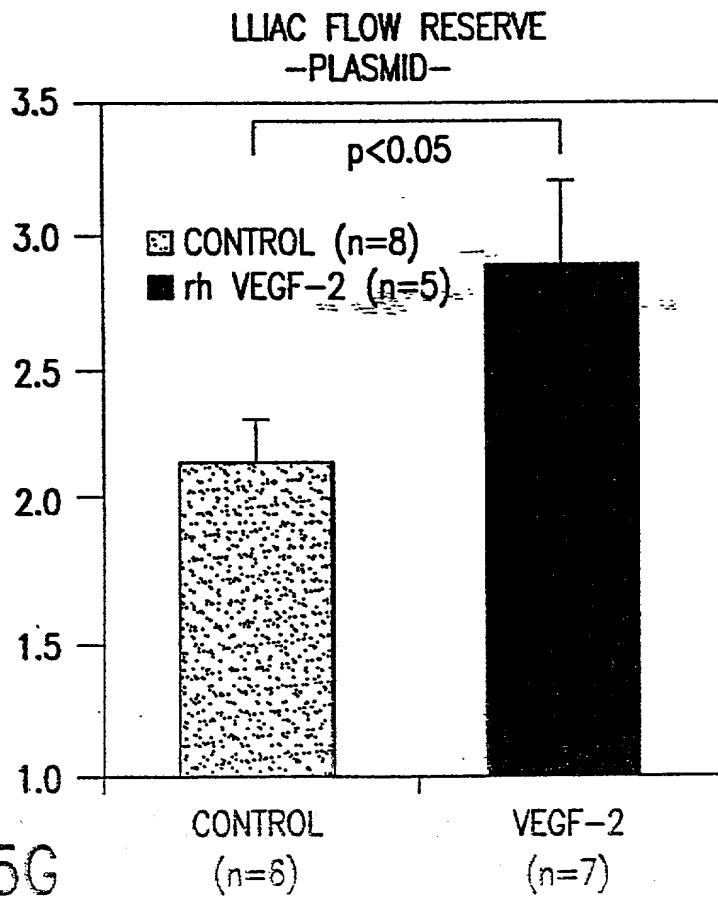


FIG.25G

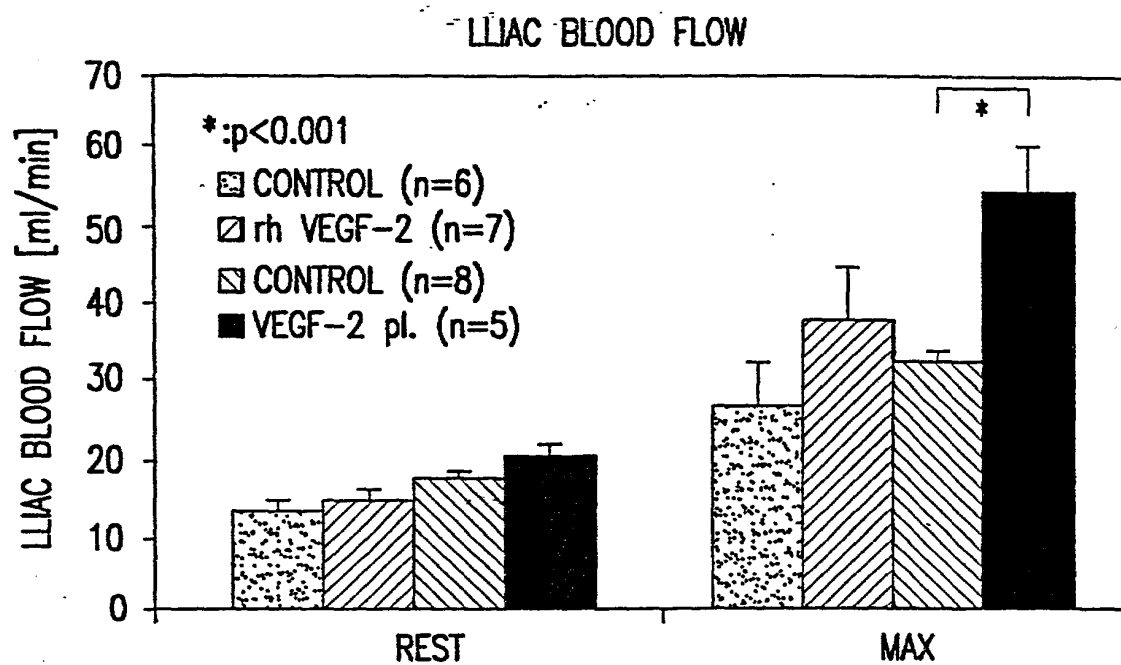


FIG.25H

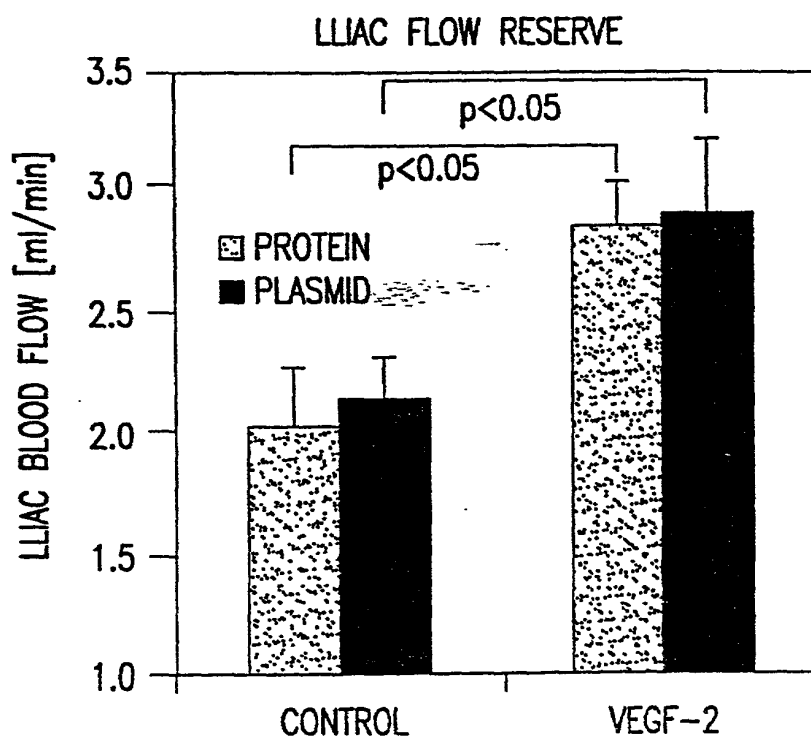


FIG.25I

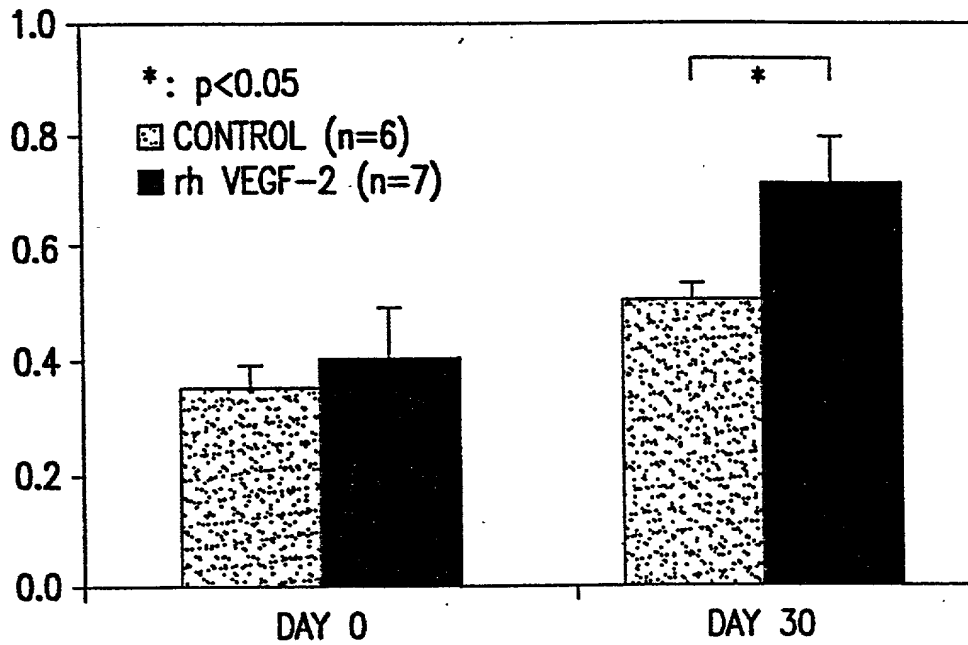
ANGIOGRAPHIC SCORE
-PROTEIN I.A.-

FIG.25J

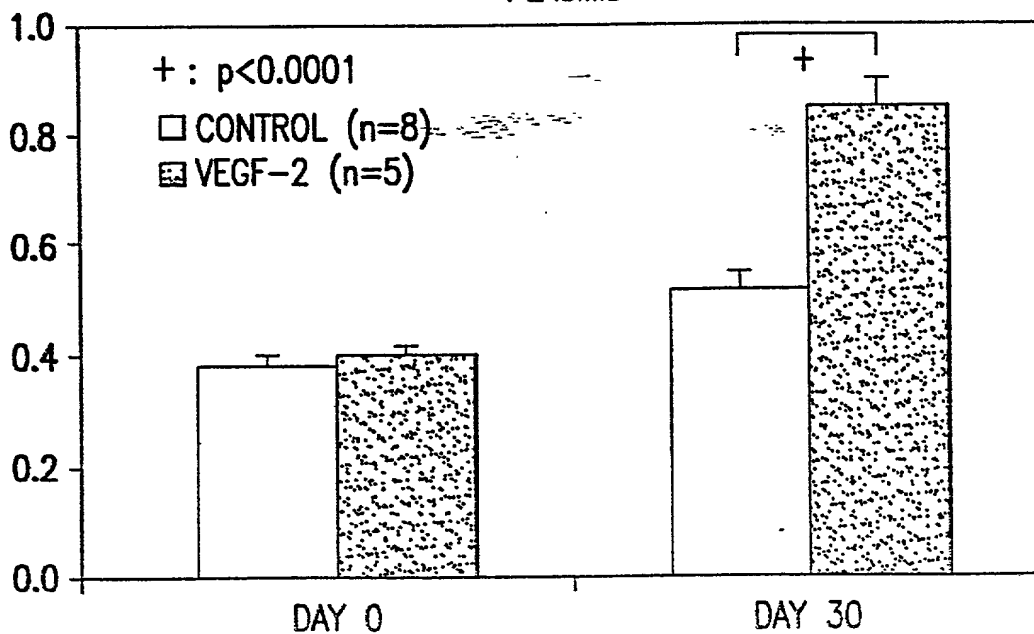
ANGIOGRAPHIC SCORE
-PLASMID-

FIG.25K

ANGIOGRAPHIC SCORE

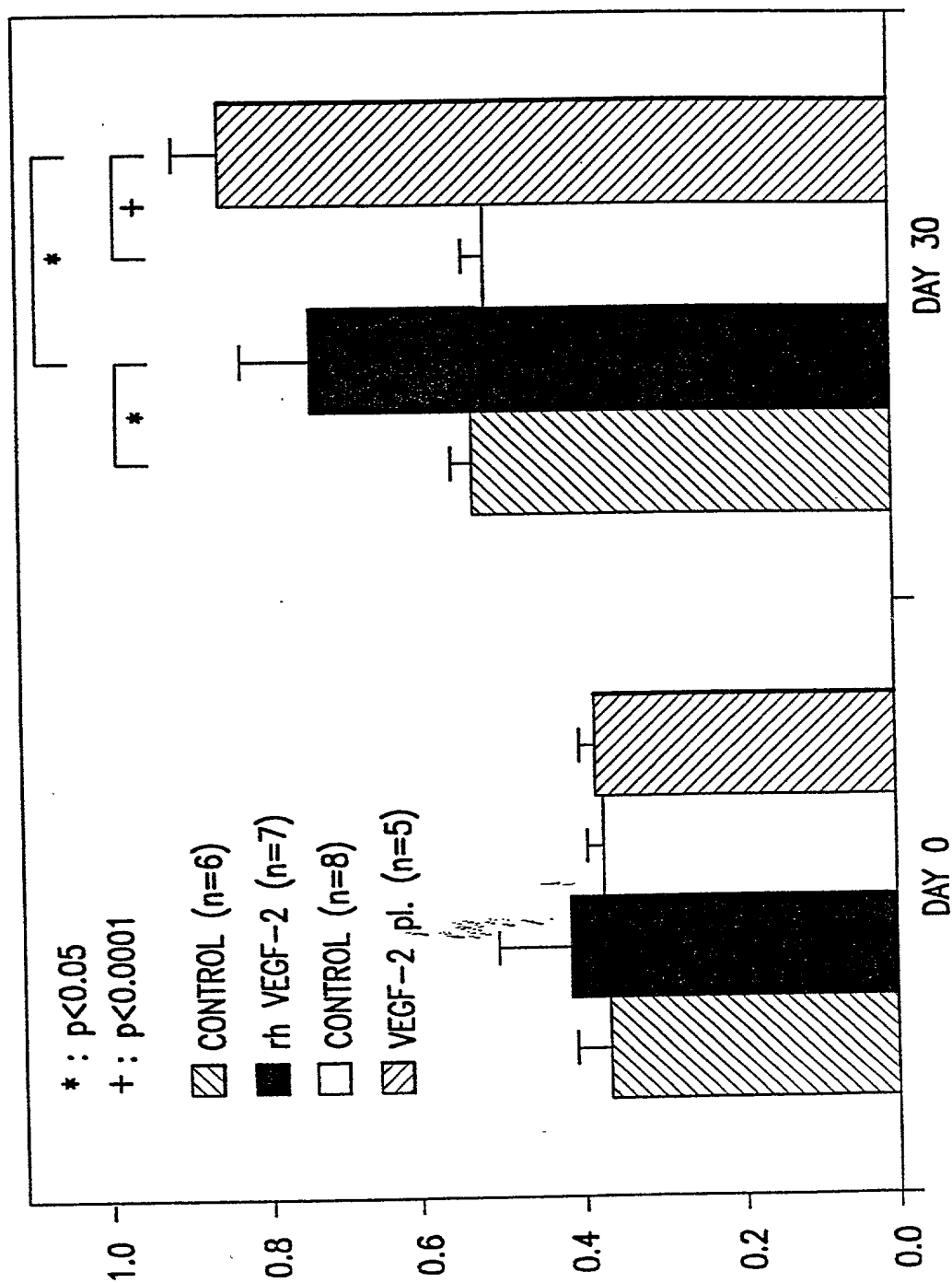
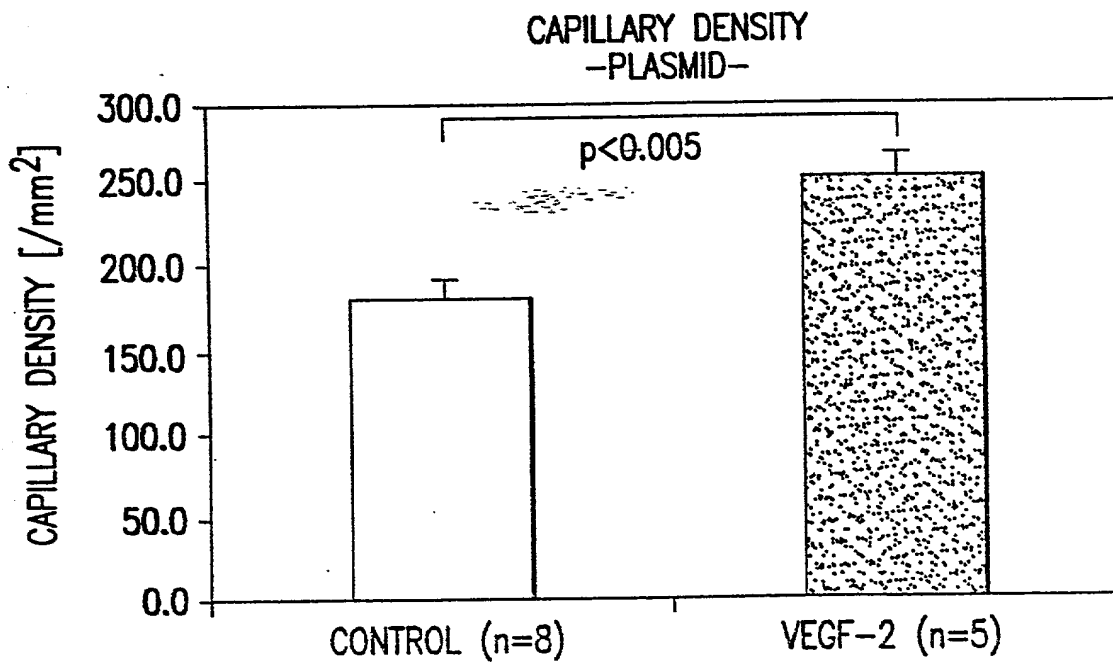
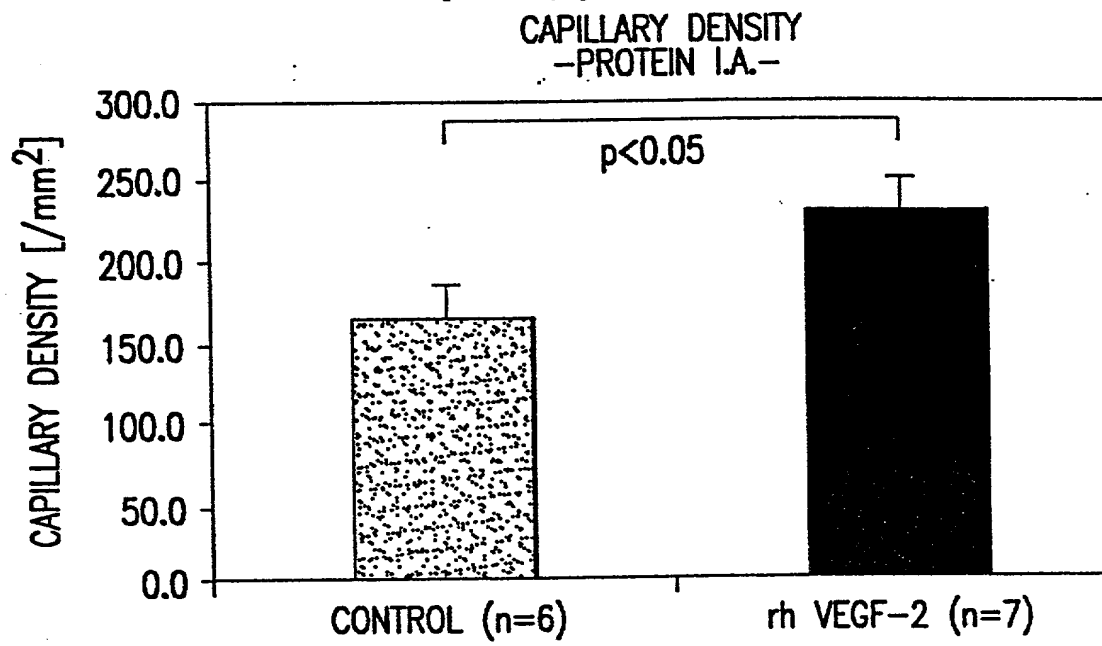


FIG.25L



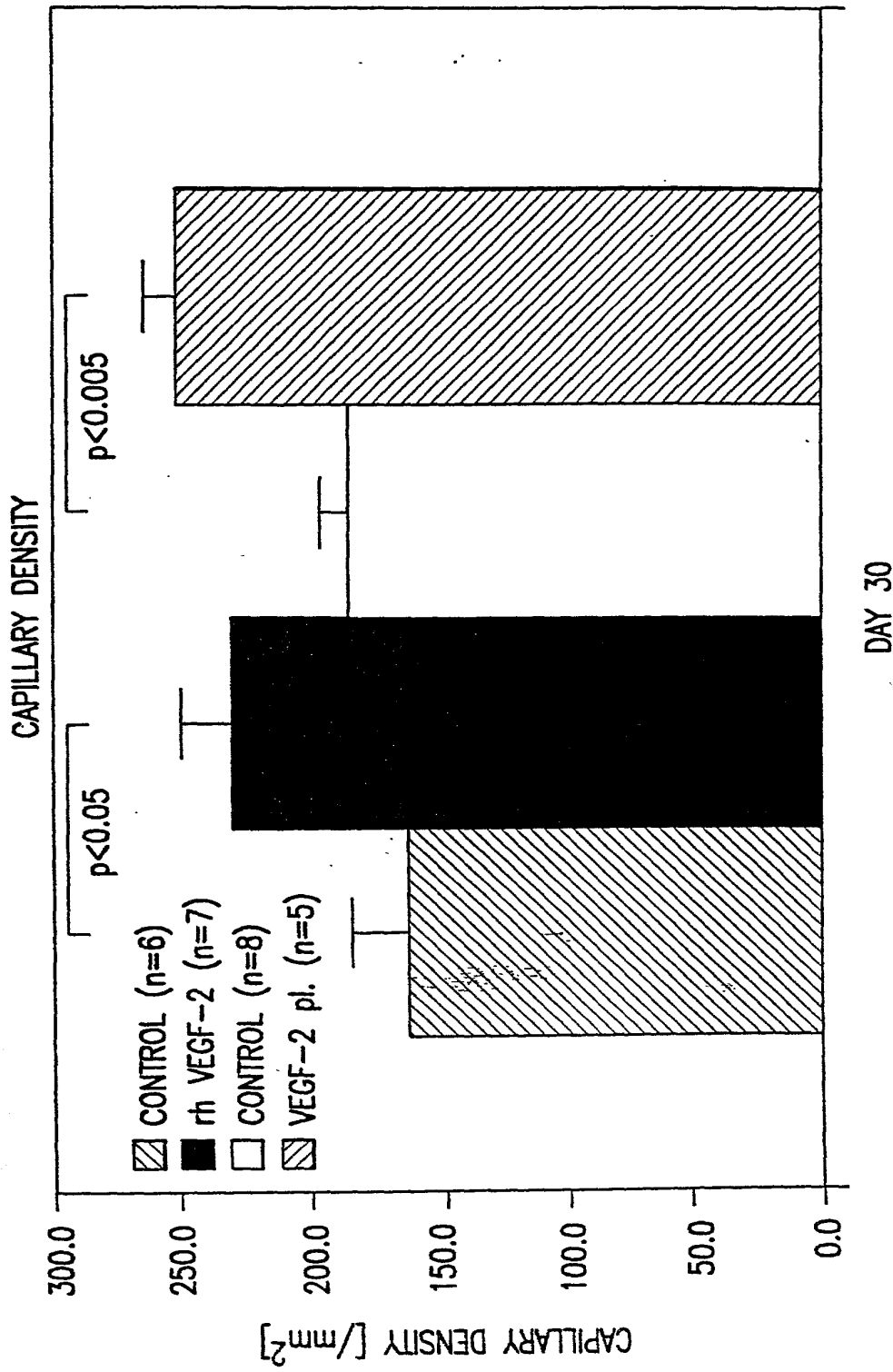


FIG.25 0

106080-4412550

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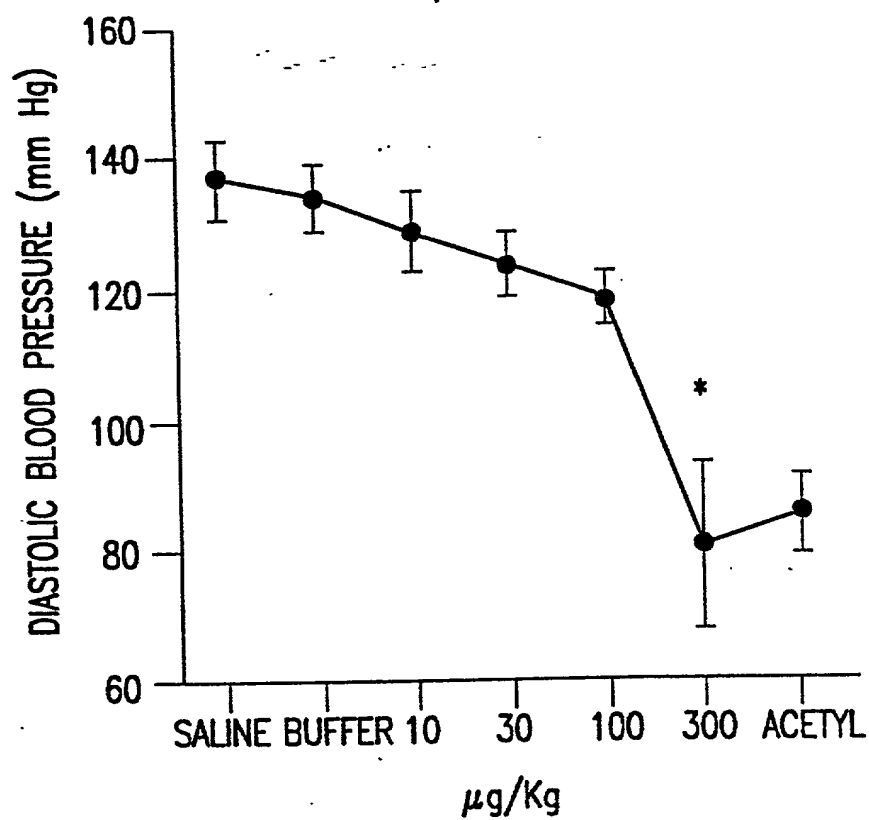


FIG.26A

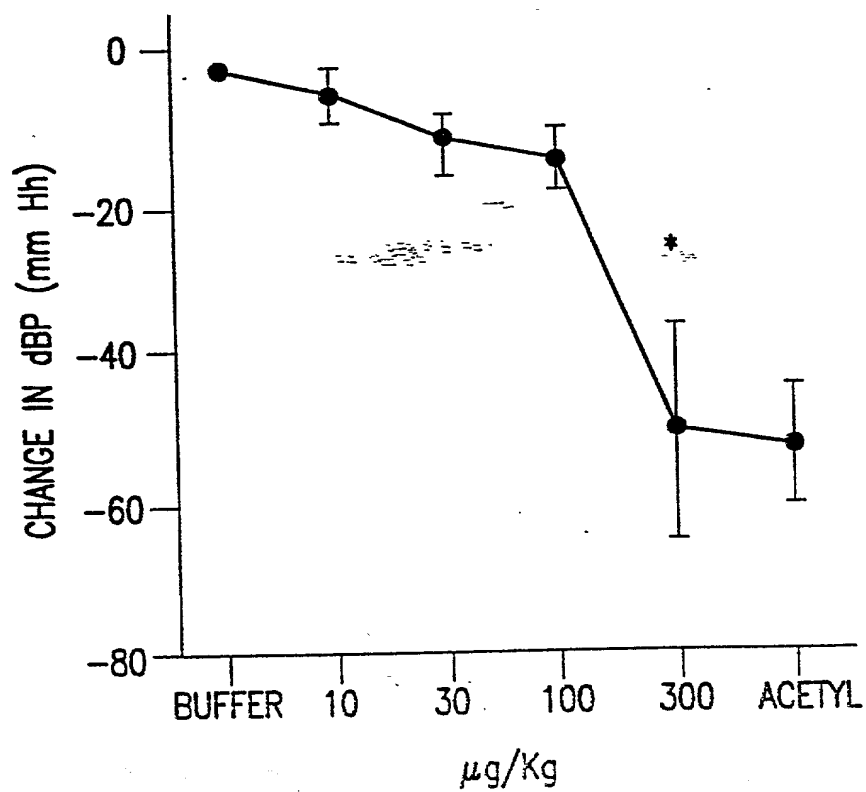


FIG.26B

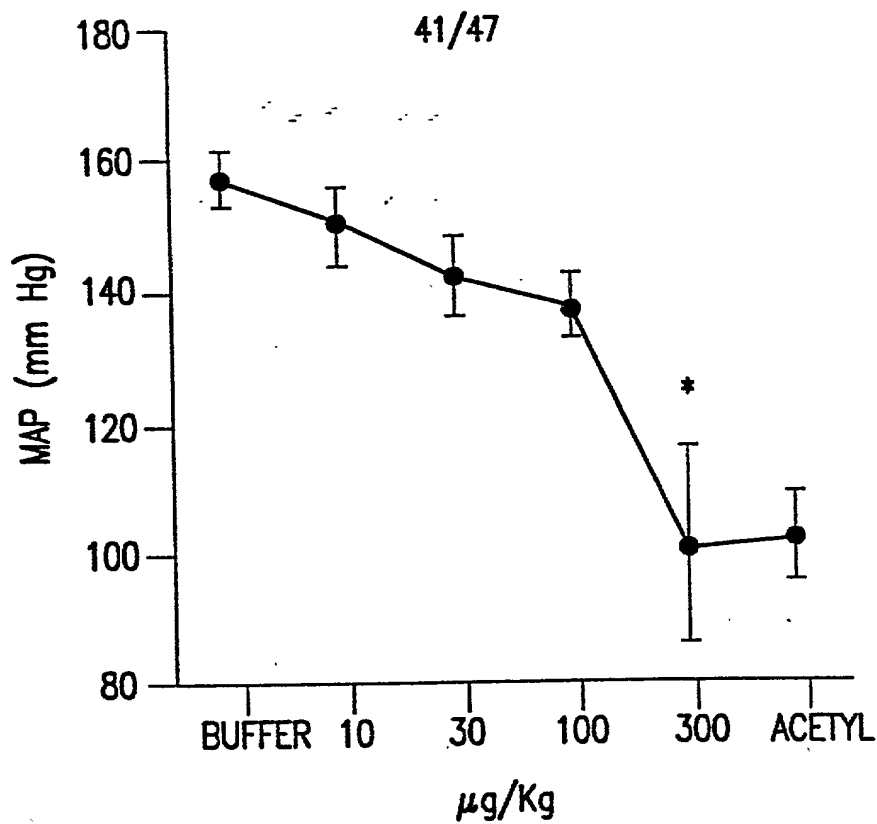


FIG.26C

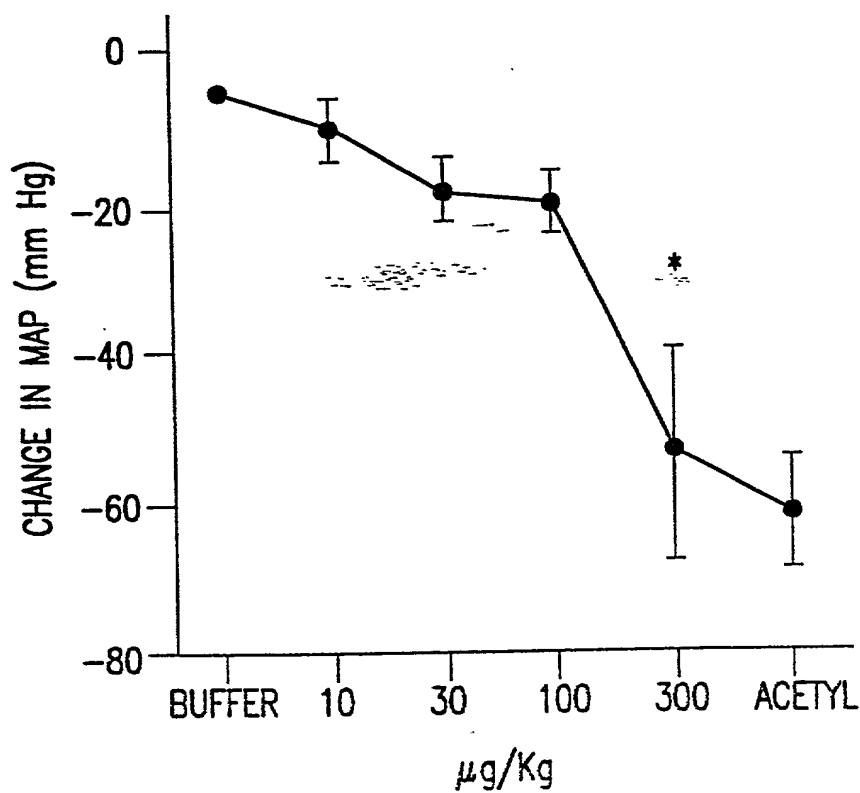


FIG.26D

CHANGE IN DBP (mm Hg)

CHANGE IN DIASTOLIC BLOOD PRESSURE OF SHR RATS GIVEN
INCREASING DOSES OF VEGF-2

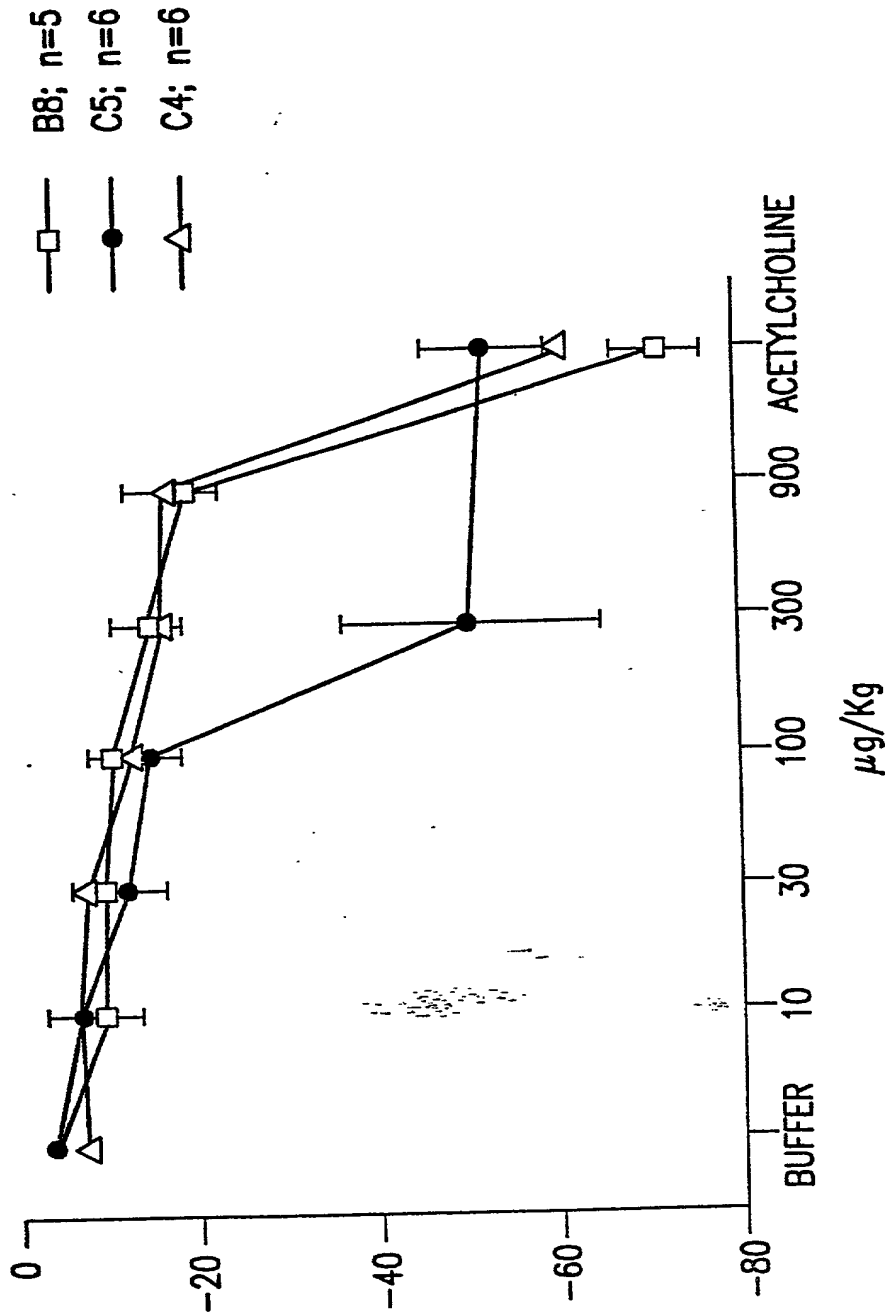


FIG.26E

THE EFFECT OF INCREASING DOSES OF VEGF-2 ON THE
MEAN ARTERIAL PRESSURE (MAP) OF SHR RATS

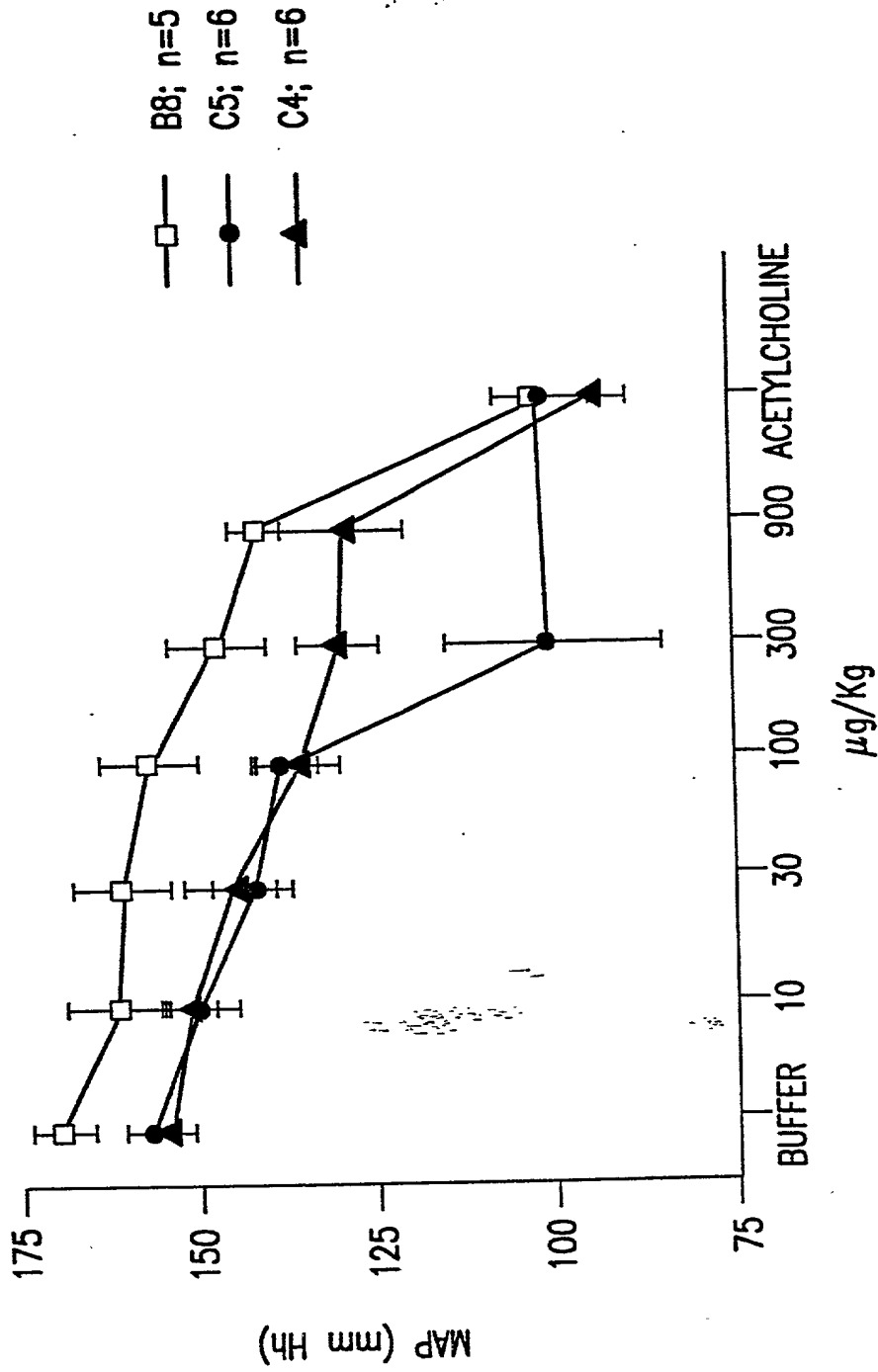


FIG.26F

THE EFFECT OF VEGF-2 ON THE DIASTOLIC BLOOD PRESSURE OF SHR RATS

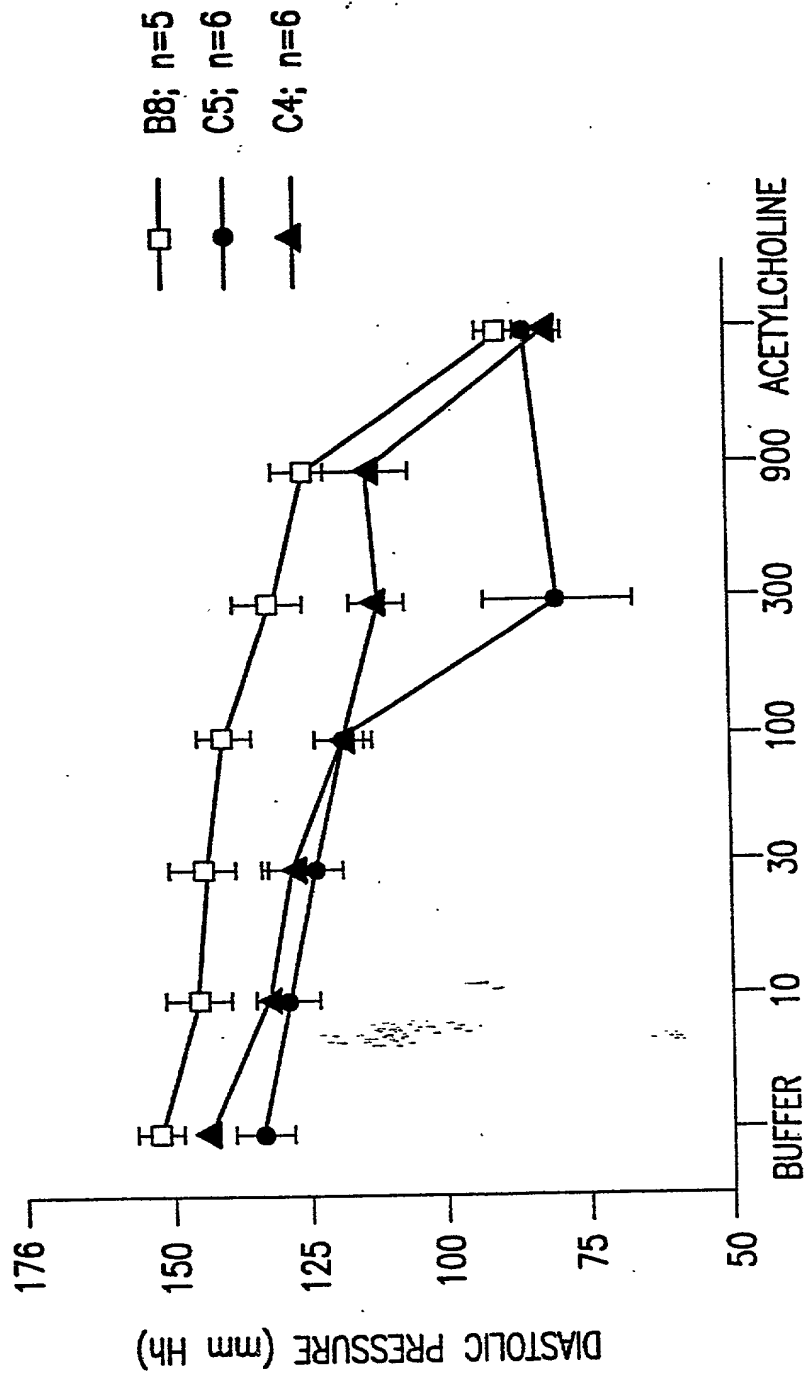


FIG.26G

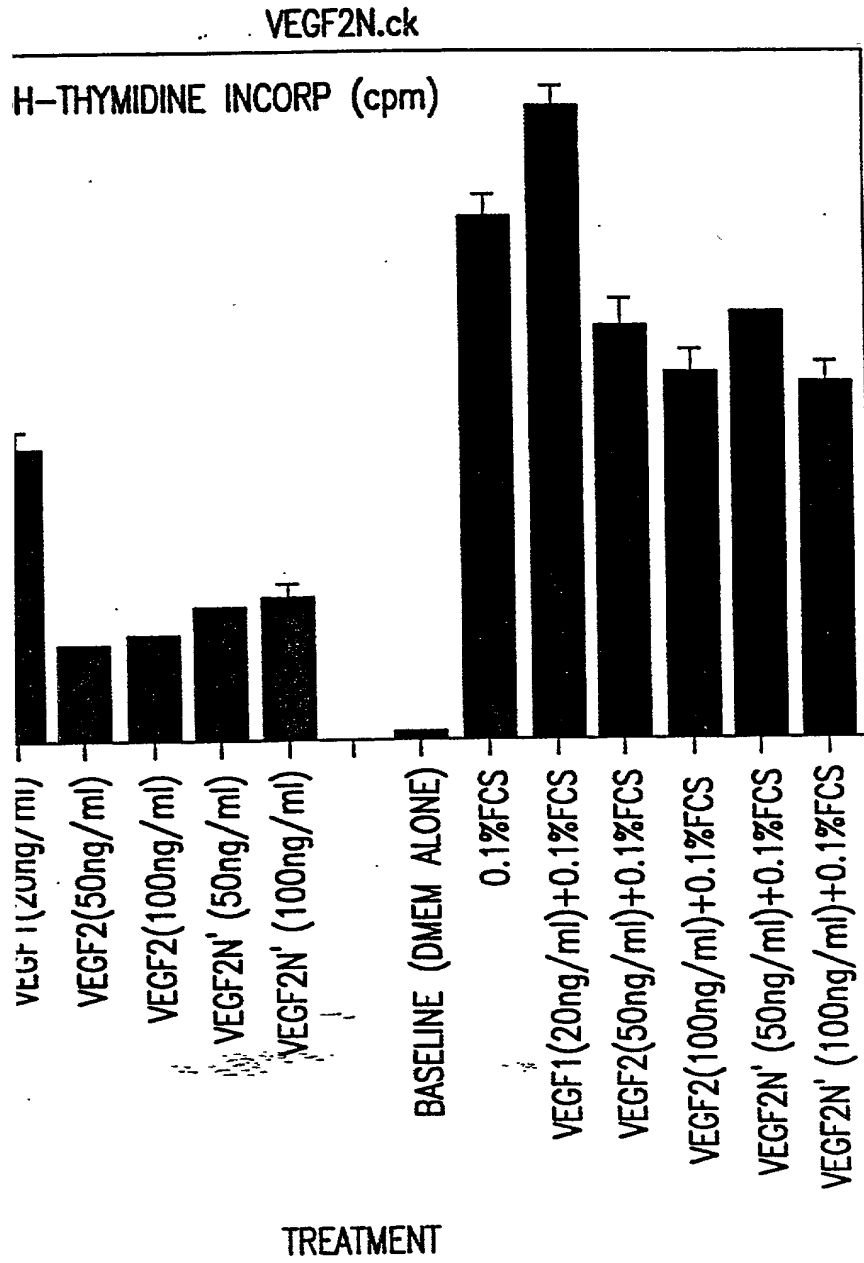
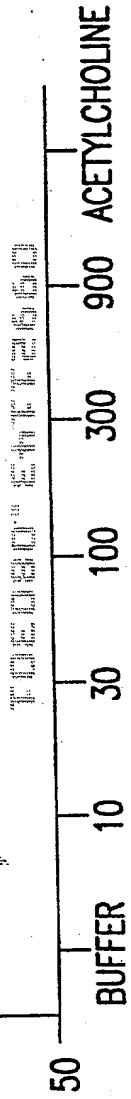


FIG.27

FIG.26G



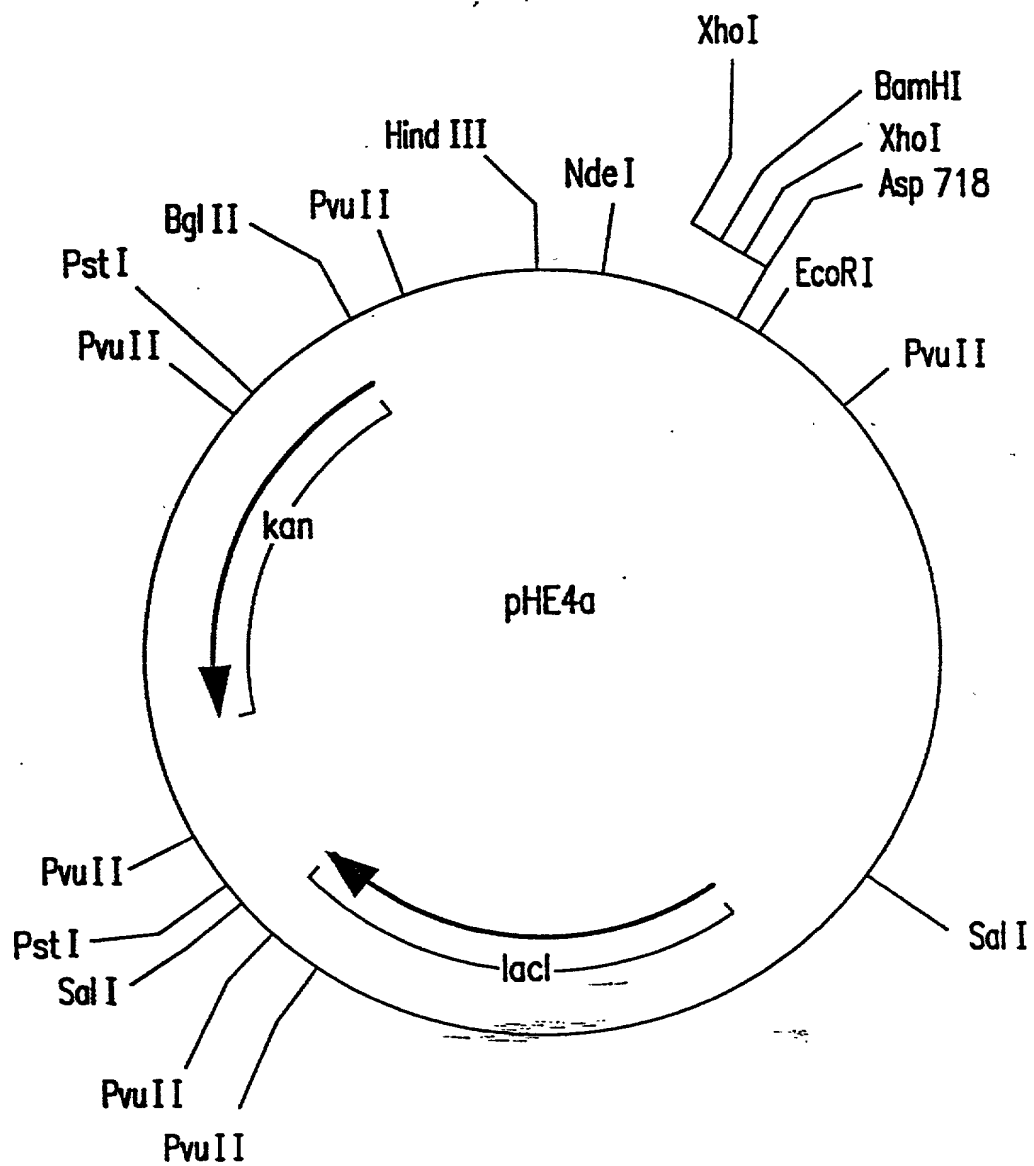


FIG.28

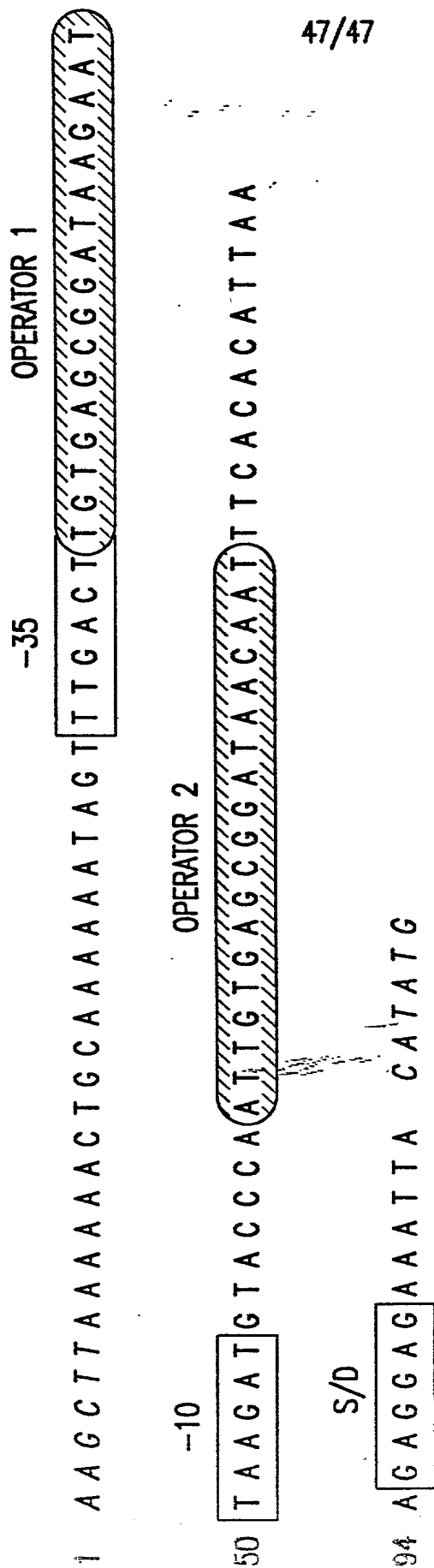


FIG.29

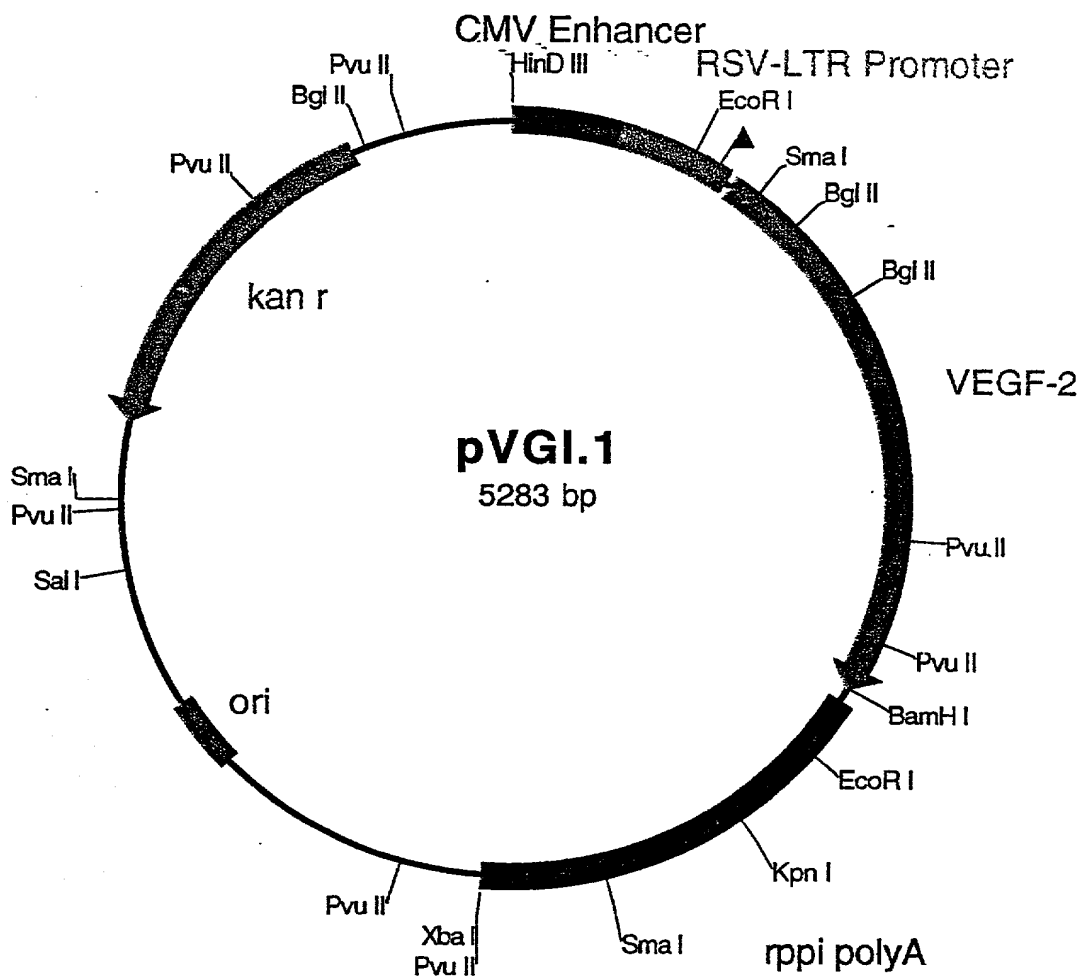


FIG. 30

FIG. 31A

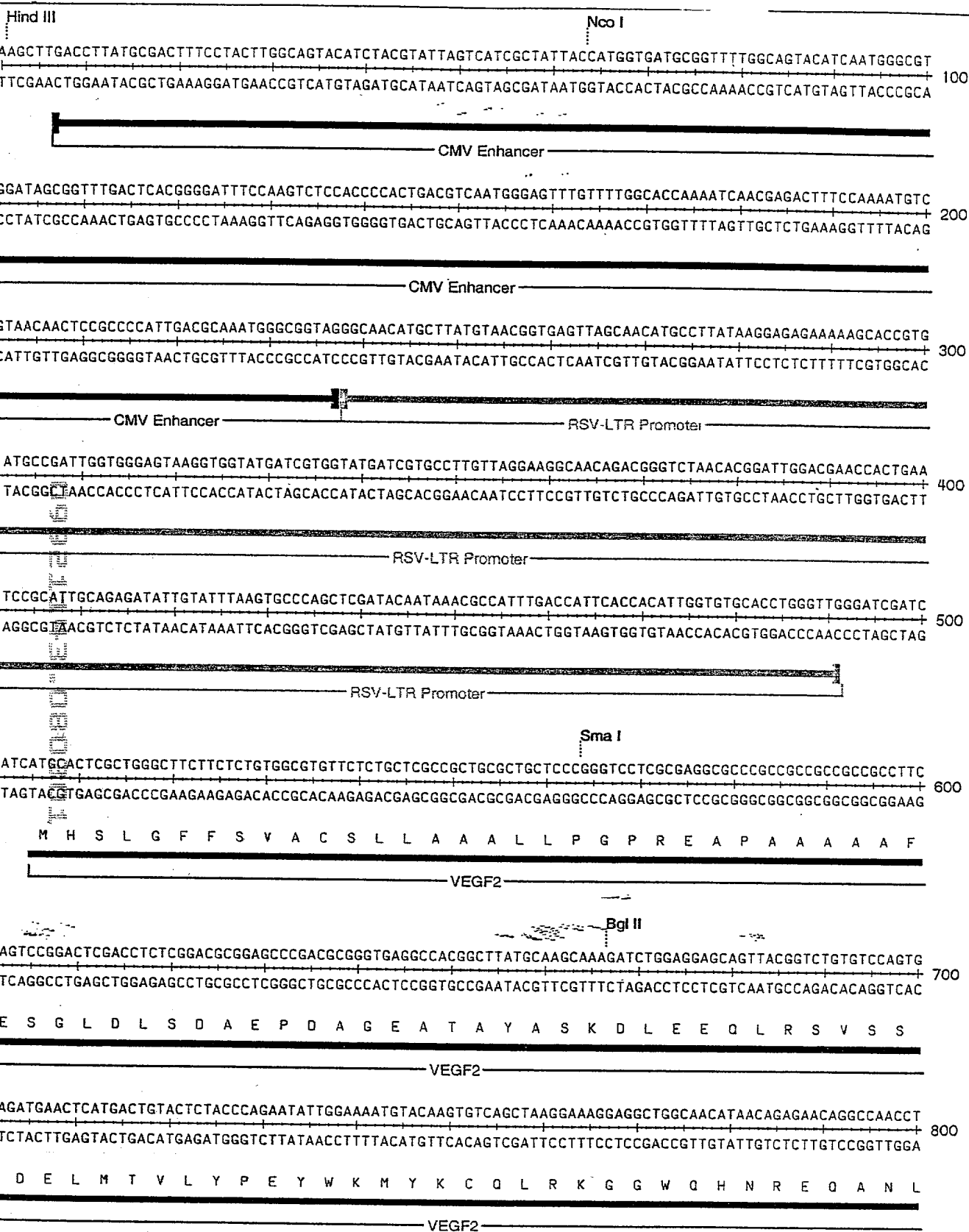


FIG. 31B

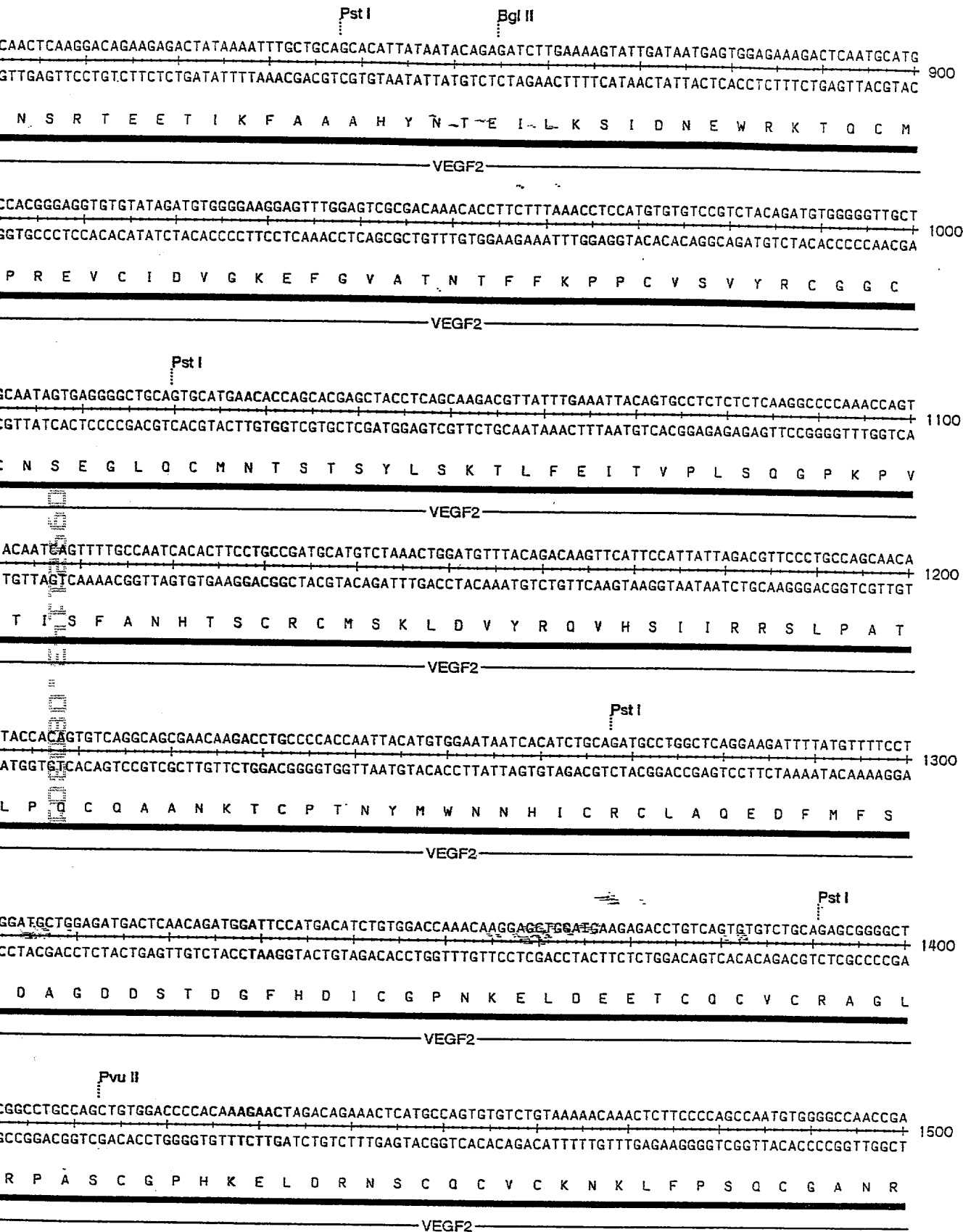


FIG. 31C

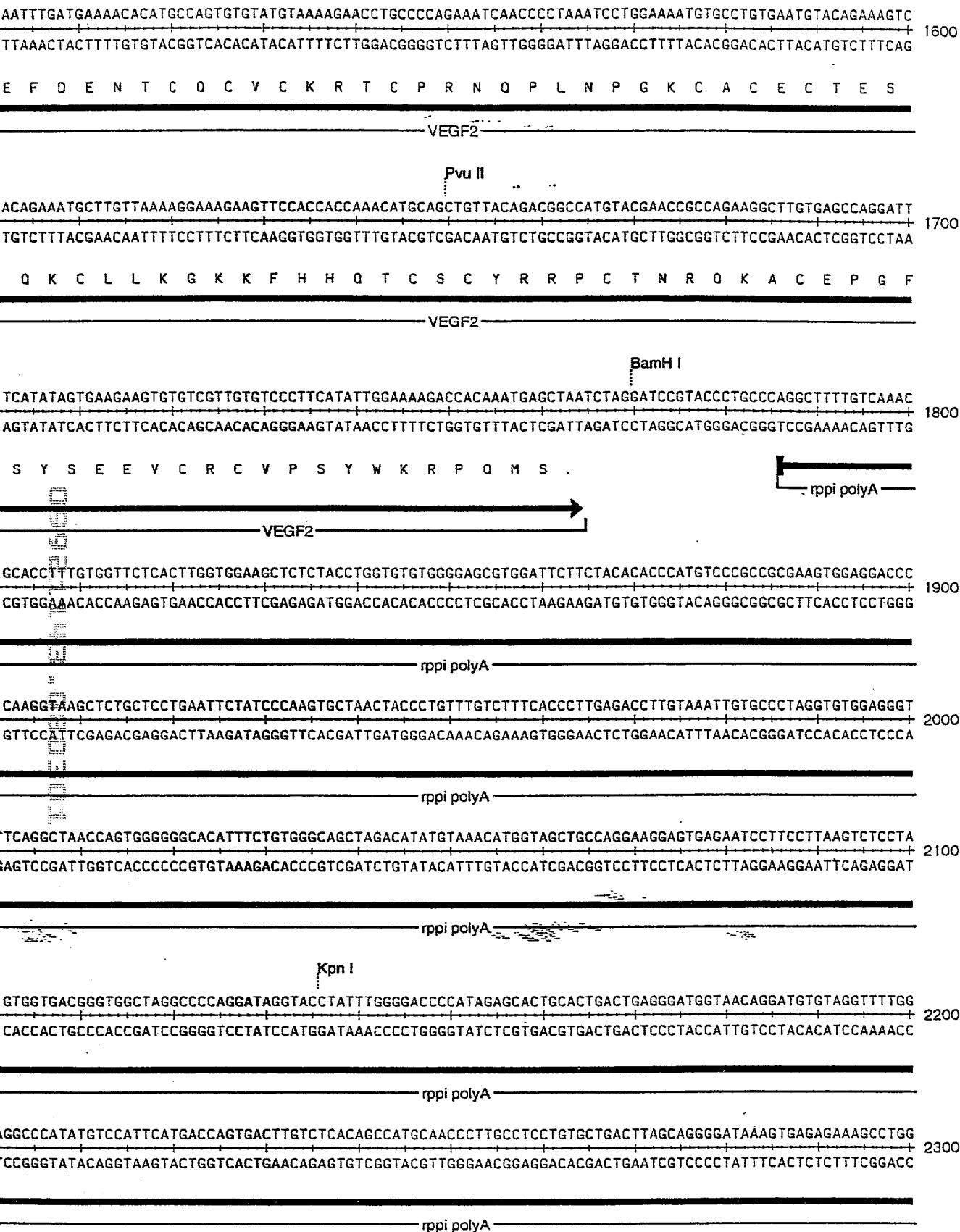


FIG. 31D

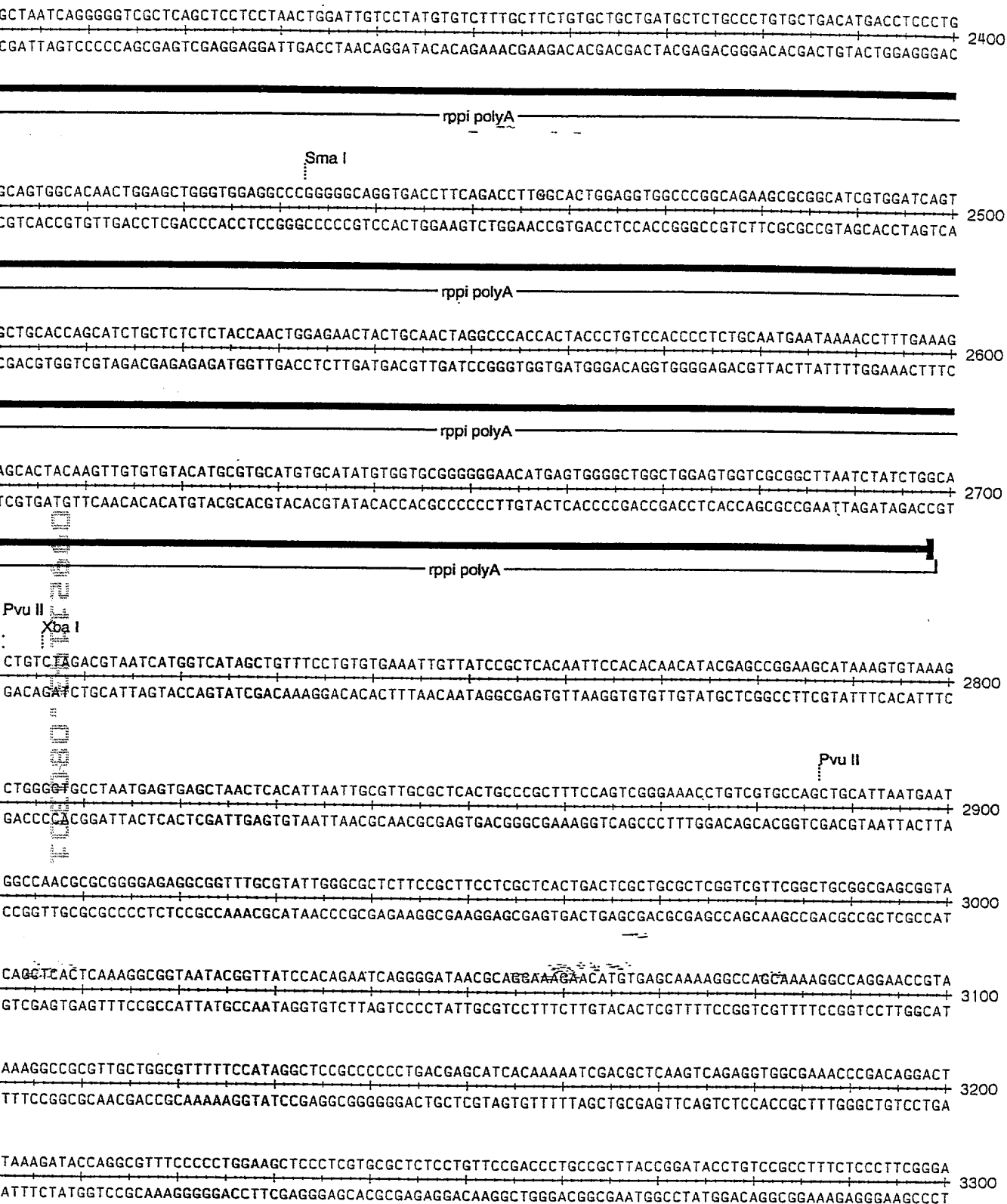


FIG. 31E

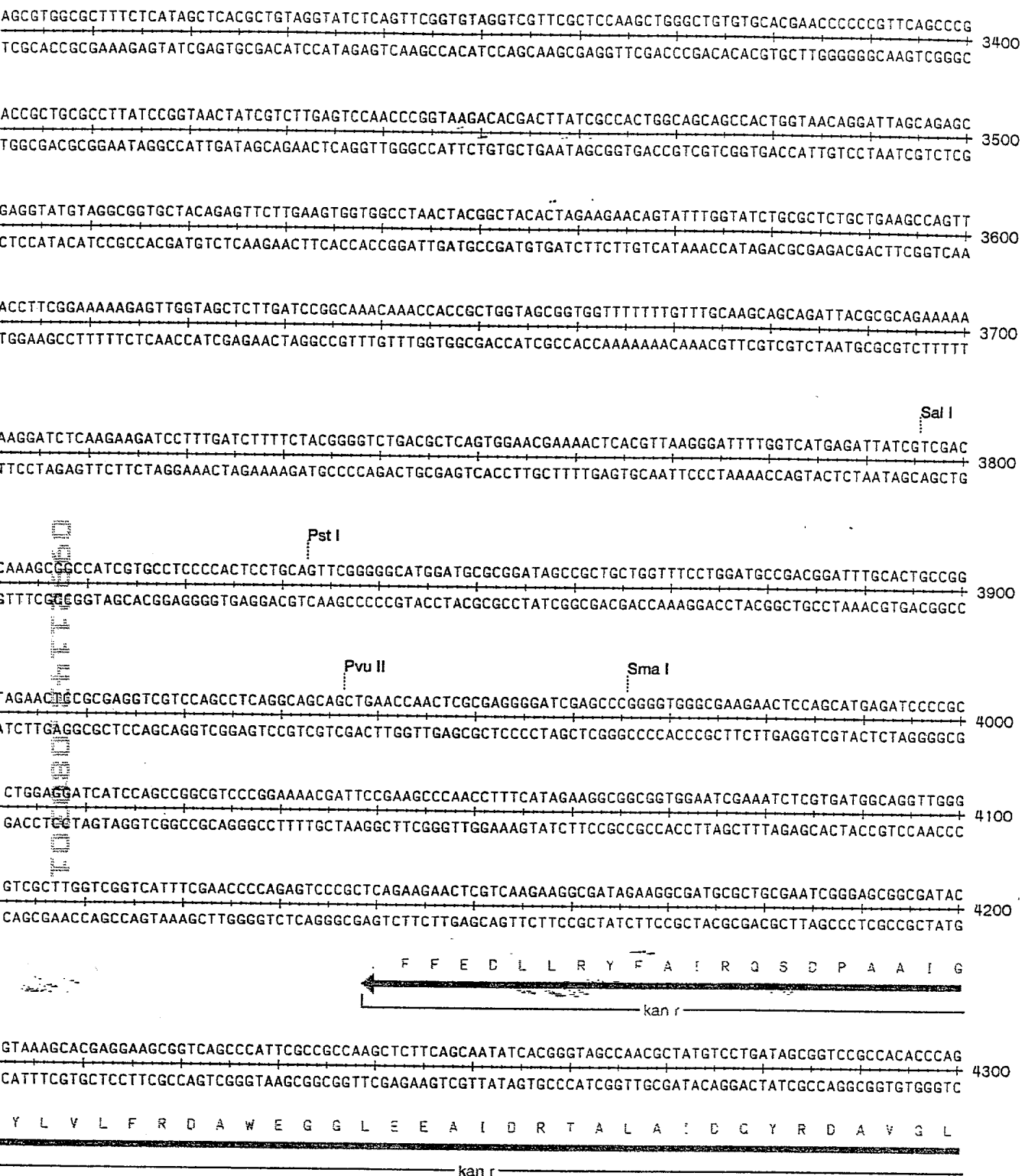


FIG. 31F

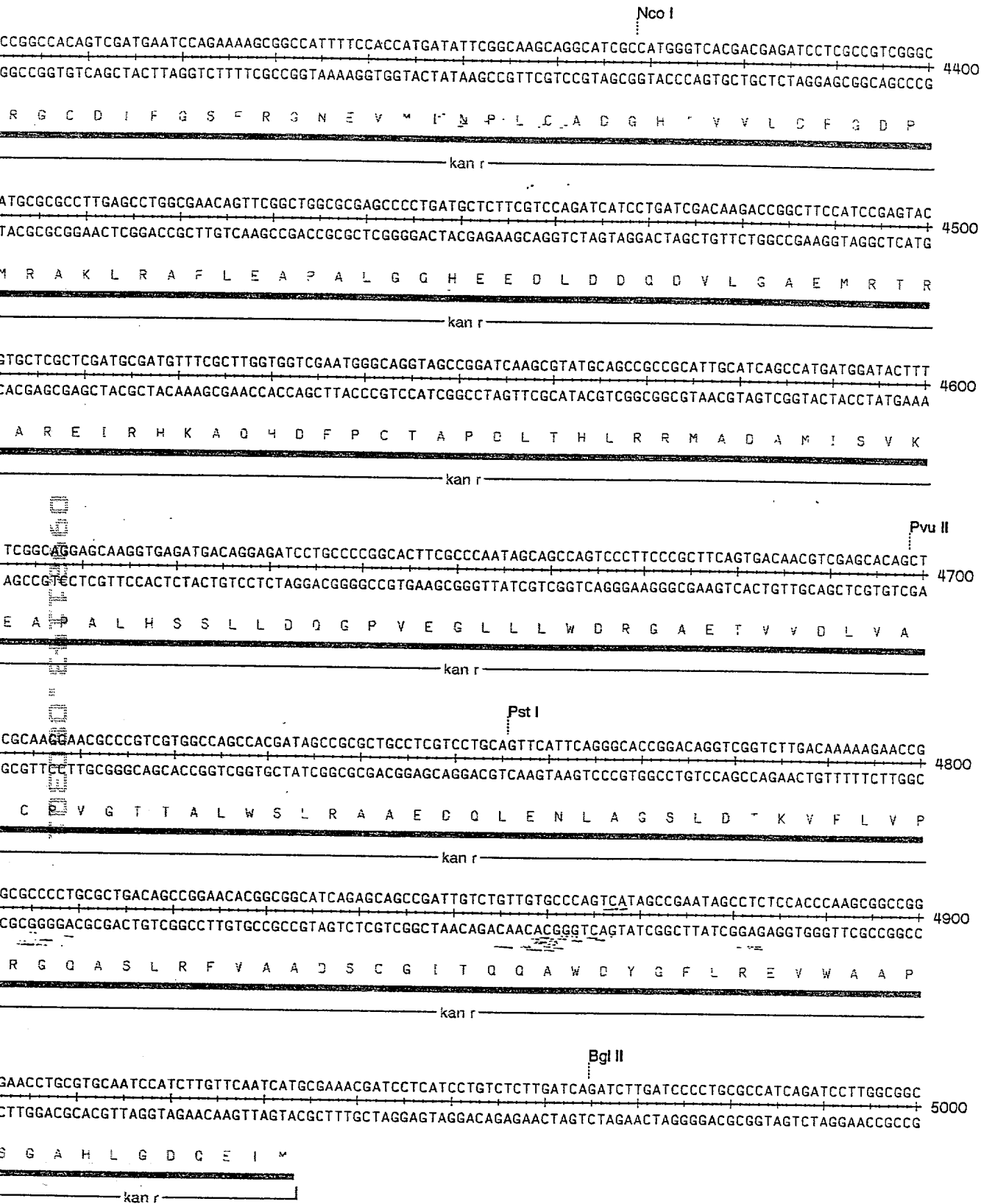


FIG. 31 G

Pvu II

AAGAAAGCCATCCAGTTTACTTTGCAGGGCTTCCCAACCTTACCAGAGGGCGCCCACTGGCAATTCCGGTTCGCTTGCTGTCCATAAAACCGCCAGT
 TTCTTTCGGTAGGTCAAATGAAACGTCCCGAAGGGTTGGAATGGTCTCCCGGGGTCGACCGTTAAGGCCAAGCGAACGACAGGTATTTGGCGGGTCA 5100
 CTAGCTATCGCCATGTAAGCCCACTGCAAGCTACCTGCTTTCTTTGCGCTTGCCTTTCCCTTGCCAGATAGCCCACTAGCTGACATTCATCCGGGG
 GATCGATAGCGGTACATTCGGGTGACGTTTCGATGGACGAAAGAGAAACGCGAACGCAAAAGGGAACAGGTCTATCGGGTCATCGACTGTAAGTAGGCCCC 5200
 TCAGCACCCTTTCTGCGGACTGGCTTTCTACGTGTTCCGCTTCCTTTAGCAGCCCTTGCGCCCTGAGTGCTTGCGGCAGCGTG
 AGTCGTGGCAAAGACGCCTGACCGAAAGATGCACAAGGCGAAGGAAATCGTCGGGAACGCGGGACTCACGAACGCCGTGCGCAC 5283

[illegible]